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NUCLEAR WEAPON EFFECT RESEARCH AT PSR—1983
Volume 10—Symptomatology of Acute Radiation Effects in Humans
after Exposure to Doses of 75 to 4500 Rads (cGy) Free-In-Air

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| 19 ABSTRACT (Continue on reverse if necessary and identify by block number) This report distills from available data descriptions of typical human symptoms in reaction to prompt ionizing radiation in the dose range 75 to 4500 rads (cgy) free-in-air. The descriptions correlate symptoms with dose and time over the acute postexposure period of six weeks. Their purpose is to provide an empirical base for estimating combat troop performance after a nuclear weapon attack. We divide the dose range of interest into eight subranges associated with important pathophysiological events. For each subrange, we estimate the signs and symptoms manifested by an exposed population—symptom onset, severity, duration, and incidence. The early or prodromal phase of radiation sickness begins about 2 to 4 hrs after doses of 300 to 530 rads (cgy). Onset time diminishes with dose, occurring within minutes of exposure to 4500 rads (cgy). Characteristic prodromal symptoms are nausea, vomiting, anorexia, and diarrhea. The prodromal phase lasts from several days to a matter of hours, depending on dose. | | | |
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19. ABSTRACT (Continued)

The delayed or manifest-illness phase begins weeks to days after exposure, onset time diminishing with increasing dose. Symptoms result primarily from injury to the hemopoietic system at doses of 150 to 1500 rads (cGy), and injury to the gastrointestinal system at doses above 1500 rads (cGy). Compound effects from both syndromes are manifested at doses of 830 to 1500 rads (cGy). Symptoms of the hemopoietic syndrome are bleeding, fever, infection, and ulceration. Symptoms of the gastrointestinal syndrome are fluid loss, electrolyte imbalance, severe diarrhea, and septicemia.

Despite differences of population characteristics, environmental conditions, and medical attention between the exposed persons represented by our data and battlefield soldiers, we believe these symptom descriptions are relevant to combat personnel.

SUMMARY

As a first step toward estimating combat troop performance after the detonation of nuclear weapons, this report describes typical human symptoms in response to prompt ionizing radiation during the acute period of six weeks after exposure.

Data on human radiation sickness symptoms are both diverse and sparse. They consist of (1) case studies of the victims of nuclear reactor accidents, (2) records of patients given radiation therapy for cancer and other diseases, (3) analyses of "composite" data, including the experience of Japanese atomic bomb survivors, and (4) expert opinion. Rather than restating the well-known variability of radiation sickness symptoms, the literature has been analyzed to reach a consensus about *typical* symptoms.

The intermediate dose range of interest, 75 to 4500 rads (cGy) free-in-air, was divided into eight subranges associated with important pathophysiological events. For each subrange, the signs and symptoms manifested by an exposed population--their onset, severity, duration, and incidence--were estimated.

The acute period of radiation response has two pathophysiological phases: an early prodromal phase and later manifest-illness phase. The prodromal phase begins about 2 to 4 hr after doses of 300 to 530 rads (cGy), and earlier with increasing dose down to minutes after exposure to 4500 rads (cGy). The characteristic signs and symptoms are nausea, vomiting, anorexia, and to a lesser degree diarrhea. Beginning at doses of about 530 rads (cGy), and as vomiting and diarrhea become severe, fluid loss, electrolyte imbalance, and headache are manifested. The prodromal phase lasts from several days to a matter of hours, depending on the dose.

The manifest-illness phase begins weeks to days after exposure, onset time diminishing with increasing dose. Symptoms result primarily from injury to the hemopoietic system at doses of 150 to 1500 rads (cGy) and injury to the gastrointestinal system at doses above

1500 rads (cGy). Compound effects from both syndromes are manifested at doses of 830 to 1500 rads (cGy). Symptoms of the hemopoietic syndrome are related to bleeding and infections. They include easy bruising, fever, and ulceration of the mouth and throat. Systemic infections are triggered by the escape of enteric bacteria from the damaged gastrointestinal mucosa; they progress fast because white blood cell production is suppressed by the radiation-caused destruction of bone marrow stem cells. Preexisting infections, for example in the respiratory tract, can rapidly become lethal. With increasing dose, and especially above 1300 rads (cGy), fluid loss and electrolyte imbalance from vomiting and injury to the gastrointestinal tract may lead to fainting, prostration, and a shock condition that could result in death in ~2 to 12 days.

Application of these descriptions to battlefield soldiers carries some reservations because of differences in population characteristics, environmental conditions, and medical attention. Accident victims and therapy patients had the benefit of medical care, which battlefield soldiers may not have. On the other hand, soldiers would presumably have advantages of youth, vigor, and motivation over the other groups mentioned. The data do not permit a quantitative assessment of the tradeoffs between postexposure medical care and preexposure robustness. We believe, however, that preexposure health condition is less important than postexposure medical care, barring prior bacterial or viral infection. The symptom descriptions may be somewhat less applicable but still reasonably relevant to combat personnel receiving doses at the lower end of the 75 to 4500 rad (cGy) range or soon after exposure. The descriptions become increasingly applicable as doses and post-exposure time increase.

PREFACE

This report was prepared as one volume of a set comprising the Pacific-Sierra Research Corporation (PSR) final report for the Defense Nuclear Agency (DNA) under contract DNA001-83-C-0015. The work done under this contract spans a wide range of nuclear weapon effects research covering intermediate-dose radiation, cratering, fire research, analytical models, underground testing instrumentation, and microwave energy. This report distills from available data descriptions of typical human symptoms in reaction to prompt ionizing radiation in the dose range 75 to 4500 rads (cGy) free-in-air. The descriptions correlate radiation sickness symptoms with dose and time over the acute post-exposure period of approximately six weeks.

The symptom descriptions are intended to provide a base for developing estimates of combat troop performance after the detonation of nuclear weapons. Such estimates, essential for military contingency planning, are the goal of the Intermediate Dose Program (IDP) of research sponsored by DNA. This work may also be of interest to policy officials and health care personnel responsible for civilian emergency management. Understanding of acute radiation effects aids planning for civil defense measures, medical facilities, and therapeutic procedures. DNA staff members David L. Auton and Cyrus P. Knowles supervised the research, and the IDP Core Group provided advisory support. Related reports include:

George H. Anno, Harold L. Brode, and Ruth Wash-ton-Brown, *Initial Human Response to Nuclear Radiation*, Pacific-Sierra Research Corporation, Note 477, April 1982 (subsequently published as DNA-TR-81-237 and Chap. II of PSR Report 1241).

George H. Anno, *Acute Radiation Response in Humans: Informal Comments by Physicians and Radiobiologists*, Pacific-Sierra Research Corporation, Note 492, rev. June 1983 (subsequently published as Vol. 14 of PSR Report 1317 and DNA-TR-82-179).

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* One centigray (cgy) is equal to one rad.
 † Unless stated otherwise, all dose levels are free-in-air values.
 ‡ This research excludes blast and thermal radiation, the two other causes of injury from nuclear weapon detonation.
 ** Throughout this report, "symptoms" is used to mean both subjective evidence and objective signs of radiation sickness.

According to U.S. Army criteria for the employment of combat units after a nuclear attack, a radiation dose of at least 3000 rads (cgy) * free-in-air † is required to render troops incapable of combat performance. Current scenarios suggest that for every soldier who receives an incapacitating radiation dose, another will receive a lethal but not incapacitating dose, 450 to 3000 rads (cgy). ‡ Two more soldiers will receive doses between "troop safety" and lethal levels, 50 to 450 rads (cgy). Many will show symptoms of radiation sickness and impaired ability to perform their normal combat tasks. The effectiveness of units manned by such sick and "walking dead" troops could become an important factor in the battlefield employment of nuclear weapons. With the continuing possibility that such weapons might be used, it is troubling that radiation-induced effects on combat performance remain poorly understood. This is the first report of a research program intended to improve our ability to predict the degree of functional impairment in military units exposed to ionizing radiation. ‡

For application to battlefield operations, the concern is with early radiation effects, those occurring within a few weeks of exposure. Because the effects of intermediate radiation doses are least well understood, the focus is on the dose range 75 to 4500 rads (cgy). Given the lack of empirical data relating combat effectiveness to radiation exposure levels, a reasonable approach is to examine the symptoms ** associated with radiation sickness and relate the symptoms to performance.

INTRODUCTION

SECTION I

As a first step in that effort, this report describes the "typical" human response to prompt radiation during the acute period of 6 weeks after exposure. The available data has been analyzed to correlate radiation sickness symptoms with dose levels and time--incidence, severity, and duration. Rather than restate the well-known variability of radiation sickness symptoms, a typical description was derived to provide a theoretical base for subsequent estimates of combat troop performance after the detonation of nuclear weapons.

The data sources fall into four general categories: (1) case studies of the victims of nuclear reactor accidents,^{*} (2) records of patients given total-body radiation therapy for cancer and other diseases,[†] (3) "composite" analyses based on data from a variety of sources, and (4) expert opinion, sometimes elicited in private communication. Additional information has come from survivors of the atomic bombings in Hiroshima and Nagasaki, as well as those accidentally irradiated in nuclear tests in the South Pacific.[‡] Though animal experiments provide the greatest quantity of data on radiation effects,^{**} animal data was not relied on because the link with human responses is indirect.

No one category of sources provides a comprehensive picture of the incidence, severity, and duration of radiation sickness in humans. For example, the data on atomic bomb survivors are usable for delayed hematological effects (≥1 week postexposure) but inadequate for early acute effects; because of the chaotic conditions, all records of symptoms during the first few postexposure days were constructed some time after the fact.^{††} To refine a plausible description of acute symptoms, it is necessary to use a variety of sources and carefully evaluate their data.

^{*}References 12, 21, 45, 51, 53, 56-58, 73, 74, 83, 90, 94, 96, 103-105.

[†]References 2, 9, 10, 15, 17, 19, 20, 42, 43, 66, 71, 78, 88, 91, 93, 98.

[‡]References 2, 26-29, 54, 55, 60, 63, 65, 72, 81, 82, 101, 106, 108.

^{**}References 3, 5, 11, 13, 14, 100, 111.

^{††}References 81, 82.

Application of these data to battlefield soldiers naturally carries some reservations because of obvious differences in population characteristics, environmental conditions, and medical attention. Accident victims and therapy patients all had the benefit of medical care in varying degrees. Battlefield soldiers might well have no access to such care. The data do not permit a quantitative assessment of the effects of such differences, but they are addressed qualitatively.

Section 2 explains the parameters adopted and assumptions made for deriving a consensus on typical symptoms from the data. Section 3 presents our findings.

SECTION 2

ANALYTIC CONSIDERATIONS

This section explains how the data were used to correlate symptoms of radiation sickness with dose levels and time after exposure for a typical population of victims. The effort involved identifying the specific *symptom*, estimating their *incidence* and *severity* in the population, reconciling *dose levels* reported in different units of measure, correlating *dose ranges* and *time* intervals with symptoms, and addressing the effect of different *dose rates* in the data. The period of acute illness is conventionally divided into the prodromal phase (1 to 3 days after exposure) and manifest-illness phase (1 to 6 weeks after exposure).

IDENTIFICATION OF SYMPTOMS

The main prodromal symptoms identified are nausea, vomiting, anorexia, diarrhea, fluid loss, and electrolyte imbalance. Concomitant effects, either a direct result of radiation or secondary to fluid loss, are headaches, fainting, and prostration. Other early effects with a different pathophysiological base are fatigue and weakness.

The manifest-illness phase is dominated by bleeding, fever, infection, and ulceration due to injury of the hemopoietic system. Higher doses can produce hypotension, dizziness, and disorientation. Fluid loss, electrolyte imbalance, and delayed diarrhea recur after relatively high doses because of damage to the intestines.*

SYMPTOM INCIDENCE AND SEVERITY

Apart from probit analyses[†] and information on atomic bomb survivors, quantitative data on the incidence of symptoms are scanty. Analysis has indicated substantial divergence between the Japanese survivor data and probit analysis predictions of symptom incidence.[‡] The reason for the

*Appendix A explores the pathophysiological basis of both prodromal and manifest-illness symptoms.

[†]References 61, 67.

[‡]References 9, 63.

*
References 67, 70.
+
Reference 67.

In this report radiation doses are expressed as free-in-air values, which are relevant to a radiation environment in the battlefield. However, most of the data sources, particularly the therapeutic accounts, express doses in midline tissue dose (MTD) values. For comparability in developing typical symptom descriptions, MTD values were adjusted, multiplying by 1.5 (1/0.66). The 0.66 factor suggested by Lushbaugh accounts for photon-dose attenuation from the surface of the body (free-in-air exposure dose) to the body midline.* The same multiplication factor was applied to estimate the free-in-air values where neutrons were present, roughly accounting for the combined first-collision neutron and secondary-photon absorbed dose at the center of the body.† This underestimates the free-in-air dose values converted from the accident data because of the mixed gamma and neutron radiation involved. For example, if an MTD neutron-gamma dose ratio of 1:3 (probably high for the accident data) is assumed, the underestimate is about 14 percent. Because of the broad dose ranges chosen (see below) and the uncertainty of the dose estimate in the accident data, that amount is insignificant for our

DOSE LEVELS

To express symptom severity, the terms used were "mild," "moderate," and "severe." This usage reflects common clinical distinctions, but we are unable to attach quantitative values to the terms for most of the symptoms identified above. Either the data do not permit quantification or quantification is inappropriate (for example, by what measure could one distinguish moderate from severe nausea or headaches?). In Sec. 3, quantitative links are shown where possible but the authors mainly rely on readers' qualitative understanding of the distinctions.

divergence is unclear. The probit data were used because they draw on more reliable accounts of accident victims and over 2000 therapy patients. However, even the probit data are not specifically correlated with post-exposure time, so the time dependence of symptom incidence has been approximated.

analysis. Of course, no adjustment was necessary for the mixed gamma and neutron radiation doses to which accident victims were exposed when reported as free-in-air values.

More precise estimates require detailed consideration of radiation transport and dosimetry (involving spectra, geometry, and composition), which is beyond the scope of this research. We do not explicitly account for relative biological effectiveness (RBE). The data do not justify values other than 1, particularly for acute symptoms such as upper and lower gastrointestinal distress, fatigue, and weakness.

DOSE RANGES

We subdivided the dose range of consideration, 75 to 4500 rads (cGy), into eight ranges reflecting important pathophysiological events, as shown in Table 1. Doses in the lowest range, 75 to 150 rads (cGy), cause minor acute damage to the hemopoietic system and mild prodromal effects (nausea, vomiting, anorexia) in a small number of irradiated persons.

In the dose range 150 to 300 rads (cGy), prodromal effects and injury to the hemopoietic system (primarily the bone marrow stem and precursor cells) increase significantly. However, victims will probably survive, except for the ~2 to 5 percent who will die after doses of about 300 rads (cGy).^{*} The probability of death increases at this dose range if victims are already weakened by other conditions, such as an infection. Although survival is possible within the next range, 300 to 530 rads (cGy), prodromal effects become pronounced. Victims also suffer moderate to severe damage to the bone marrow. As the dose reaches about 500 rads (cGy), 50 percent who do not receive appropriate medical care may die within 60 days.[†]

The lower limit of the 530 to 830 rad (cGy) dose range is the estimated LD_{50/60}; 100 percent lethality is approached at about 750 rads (cGy).[‡] Responses to doses between 830 and 1100 rads (cGy) begin to

^{*}References 61, 75.

[†]Reference 61.

[‡]References 61, 77.

Table 1. Dose ranges and associated pathophysiological events.

| Pathophysiological Events | | | | | | |
|------------------------------|----------------------|---|--|---------------------------|--|--|
| Dose Range, rads (cGy) | Prodromal Effects | Manifest-Illness Effects | Survival | | | |
| 75-150 | Mild | Slight decrease in blood cell count | Virtually certain | Probable (>90 percent) | | |
| 150-300 | Mild to moderate | Beginning symptoms of bone marrow damage | | | | |
| 300-530 | Moderate | Moderate to severe bone marrow damage | Possible-- Bottom third of range: LD ₅ /60 Middle third: LD ₁₀ /60 Top third: LD ₅₀ /60 | Death within 3½-6 weeks-- | | |
| 530-830 | Severe | Severe bone marrow damage | Bottom half: LD ₉₀ /60 Top half: LD ₉₉ /60 | Death within 2-3 weeks | | |
| 830-1100 | Severe | Bone marrow pan- cytopenia and moderate in- testinal damage | | Death within 1-2½ weeks | | |
| 1100-1500 | Severe | Combined gastroin- testinal and bone marrow damage; hypotension | | Death within 5-12 days | | |
| 1500-3000 | | Severe gastrointestinal damage Upper half of range: early transient incapacitation; gastrointestinal death | | Death within 2-5 days | | |
| 3000-4500 | | Gastrointestinal and cardio- vascular damage | | | | |

reflect the combined effects of gastrointestinal and hemopoietic damage. Survival is almost impossible unless a compatible bone marrow transplant is available. Nearly everyone irradiated at this level suffers severe prodromal effects during the first day after exposure.

Injuries from doses of 1100 to 1500 rads (cGy) are similar to those in the foregoing dose ranges but much more severe due to greater depletion of bone marrow stem cells,^{*} greater gastrointestinal damage, and systemic complications. Early transient incapacitation has been observed in monkeys and may appear in man.[†] At 1500 to 3000 rads (cGy) early transient incapacitation may become more frequent.[‡] An early-postexposure renal failure was reported in this dose range.^{**} Death results in less than 2 weeks from gastrointestinal injury, complicated by bone marrow damage and concomitant cessation of granulocyte production.^{††} Above about 2000 rads (cGy), death may occur sooner from electrolyte imbalance and dehydration due to vomiting and diarrhea, especially in hot or humid conditions. Extremely severe gastrointestinal and cardiovascular damage causes death within 2 to 5 days after doses of 3000 to 4500 rads (cGy).^{‡‡}

POSTEXPOSURE TIME CORRELATIONS

The onset and duration of prodromal symptoms in relation to radiation dose was estimated primarily from accounts of accident victims and therapy patients. Data on the Japanese atom bomb survivors are inadequate for prodromal symptom-time correlations, as noted in Sec. 1. For manifest-illness hemopoietic depression, however, the Japanese data are more reliable, having been recorded at the time of occurrence.^{***}

*References 7, 65.

†Reference 23.

‡Reference 23.

**Reference 98.

††Reference 65.

‡‡Reference 68.

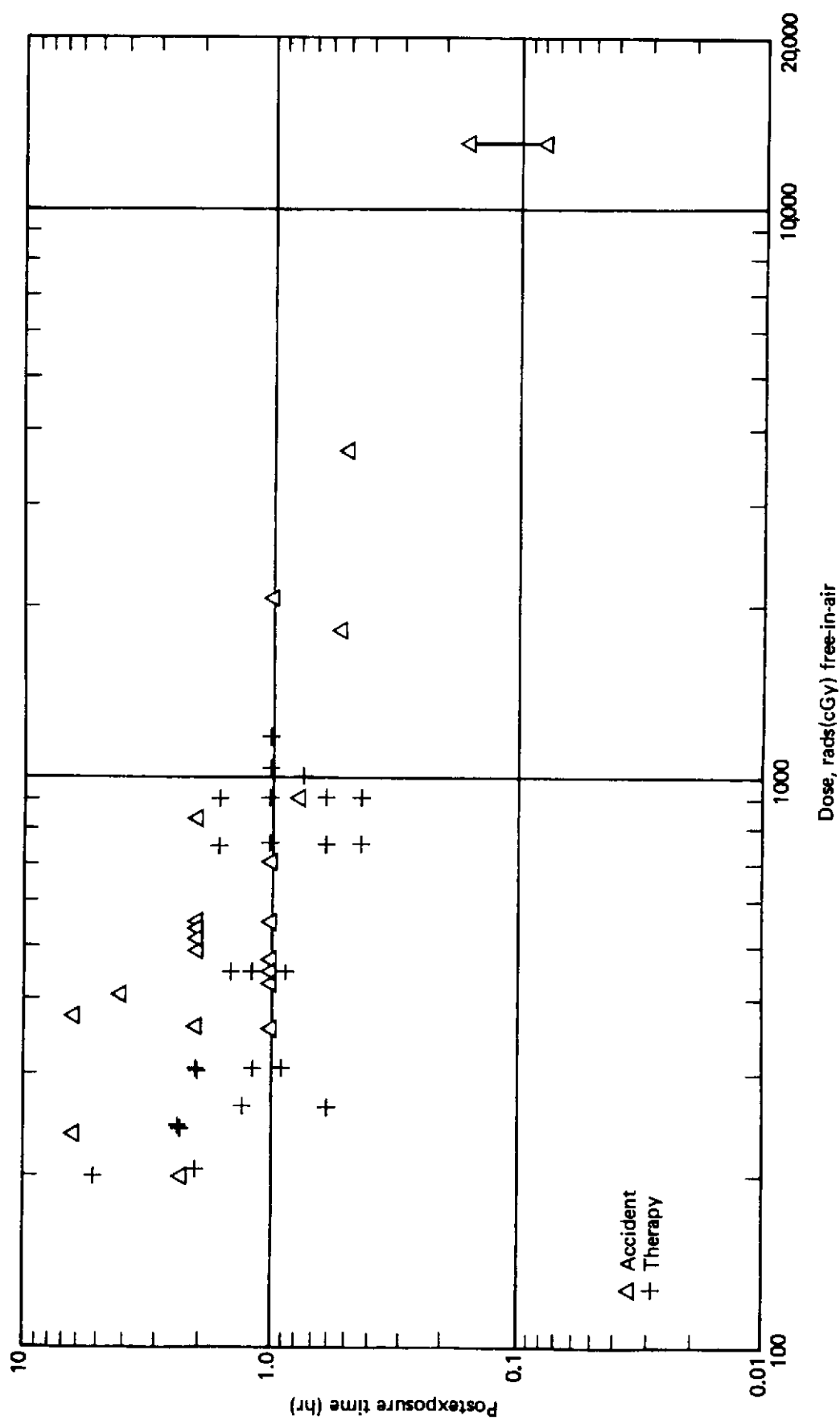
***References 2, 29, 60, 81, 82, 101, 106.

It is well known from radiobiological research that tissue cells are affected differently by different radiation dose rates as well as by different total doses. The data in this report include a great variety of dose rates. Accident victims were exposed to many thousands of rads in a fraction of a second. * In contrast, patients undergoing total-body irradiation were exposed to 1 to 30 rads (cGy)/min over periods of minutes to hours. We considered whether an adjustment of the therapy data for a dose rate differential to develop typical symptom descriptions was needed. Given the great differences in dose rate between accident victims and therapy patients, it might be expected that their prodromal symptoms would begin at different times. However, correlation of the onset time of prodromal symptoms with dose level (Fig. 1) shows no marked difference between the two groups. † As for therapy patients alone, radiation therapists and radiobiologists recently said they had found no evidence of earlier onset of worse nausea or vomiting with increasing dose rate in the therapeutic range indicated above. ‡ It is believed, therefore, that for deriving typical acute response patterns for symptoms such as nausea, vomiting, and fatigability, data from therapy patients are directly applicable.

*References 69, 70, 89.

†Reasoning from two studies of accident victims (Refs. 29 and 73), one analyst has suggested that prodromal symptoms and hematological injury are less severe with small daily doses at low dose rates than single high-intensity doses of equal size (Ref. 61). However, the dose rates for those accidents [0.05 and 0.02 rads (cGy)/min] were considerably lower than even the dose rates in our therapy patient data.

‡References 40, 99.



SECTION 3

FINDINGS

This section sets forth descriptions of the typical course of radiation sickness for each of the eight dose ranges identified in Sec. 2. The descriptions are preceded by a discussion of our findings on the onset and duration of symptoms, symptom incidence, and radiation lethality.

ONSET AND DURATION OF SYMPTOMS

The onset of symptoms was estimated by plotting the relation between the time prodromal symptoms began and the dose (see Fig. 2). Logarithmic presentation is convenient because of the large range of actual doses--190 to 13,200 rads (cGy). Data points are indicated by symbols representing the four data categories:

| Category | References |
|-----------------------|--------------------------------|
| Accident (Δ) | 30, 38, 52, 59, 62, 96, 105 |
| Therapy (+) | 17, 78, 88, 92 |
| Composite (O) | 42, 44, 46, 62, 69, 79, 110 |
| Expert opinion (x) | 9, 38, 42, 44, 46, 90, 96, 105 |

Where sources disagreed, we favored accident and therapy data over the other two categories. Each symbol indicates firm information; lines connecting the symbols indicate less certain ranges; arrows indicate open-ended values based on quite uncertain data.

Because the data vary greatly in density and precision, it was inappropriate to apply numerical techniques such as regression analysis. Instead, a curve was drawn through the data points to represent "typical" individuals. The curve shows onset time to be inversely proportional to dose.

The exact trend at the high end of the dose range is uncertain because of the lack of empirical data. However, the curve is supported by

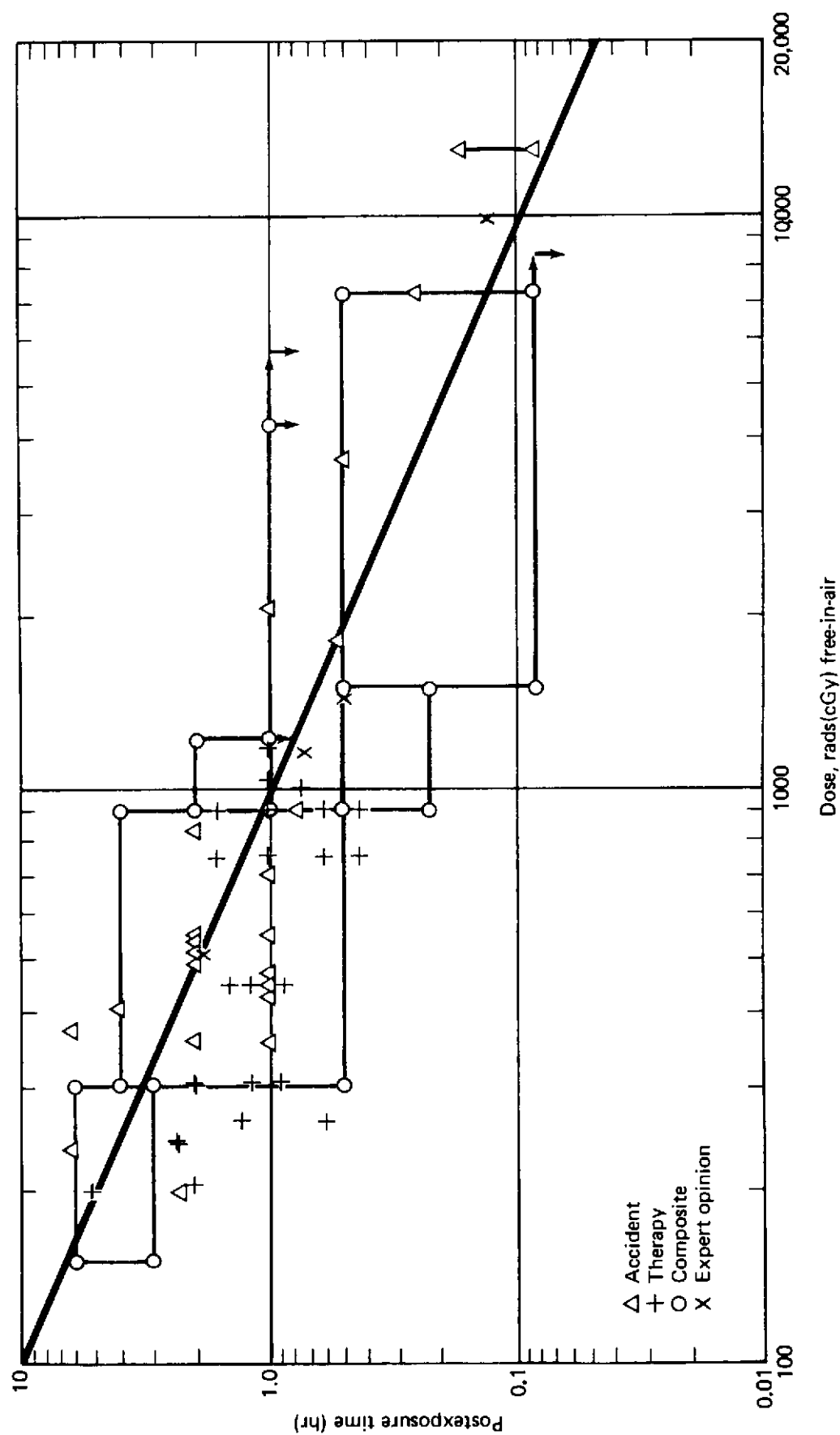


Figure 2. Onset of prodromal symptoms related to dose--all categories of data.

the accident data point at 13,200 rads (cGy) and by Langham's opinion that persons exposed to several thousand rads (MTD) will probably show the entire range of prodromal symptoms within 5 to 15 min.[†]

To check the representativeness of the curve, the temporal distribution of the onset of vomiting was examined in 100 male victims.[†] The

The mean onset time was 144 ± 66 min after exposure to single doses above 450 rads (cGy). At 450 rads (cGy) the Fig. 2 curve corresponds to an onset time of about 2 hr, reasonably close to 144 min (2.4 hr), given the imprecision of the data. Moreover, with a standard deviation of ± 66 min (1.1 hr), and assuming an approximately normal distribution, prodromal symptom onset would be expected at 0.5 to 4.3 hr for 92 percent (i.e., 1.73 σ) of those exposed. Indeed, the vomiting data and the curve are consistent with those confidence bounds.

To estimate the duration of symptoms in the prodromal and manifest-illness phases, we began with previously developed frameworks relating phase duration to dose.^{**} Deriving a consensus on symptoms from the data, we expanded the framework to specify the duration of each symptom identified in Sec. 2, related to dose.

INCIDENCE OF SYMPTOMS

Figure 3 shows the results of probit analyses relating the incidence of prodromal symptoms to radiation dose.^{††} Plotted on lognormal probability paper, the straight-line curves assume a lognormal distribution and the concave curves, a normal distribution.^{††} The lognormal distribution generally fits the data better. Exceptions are the incidence of

*Reference 59.

†Reference 61.

‡Reference 67.

**References 9, 61.

††References 61, 67, 69, 70, 105.

††Lognormal curves are based on data from accident victims and over 2000 therapy patients; normal curves are based on data from 163 therapy patients.

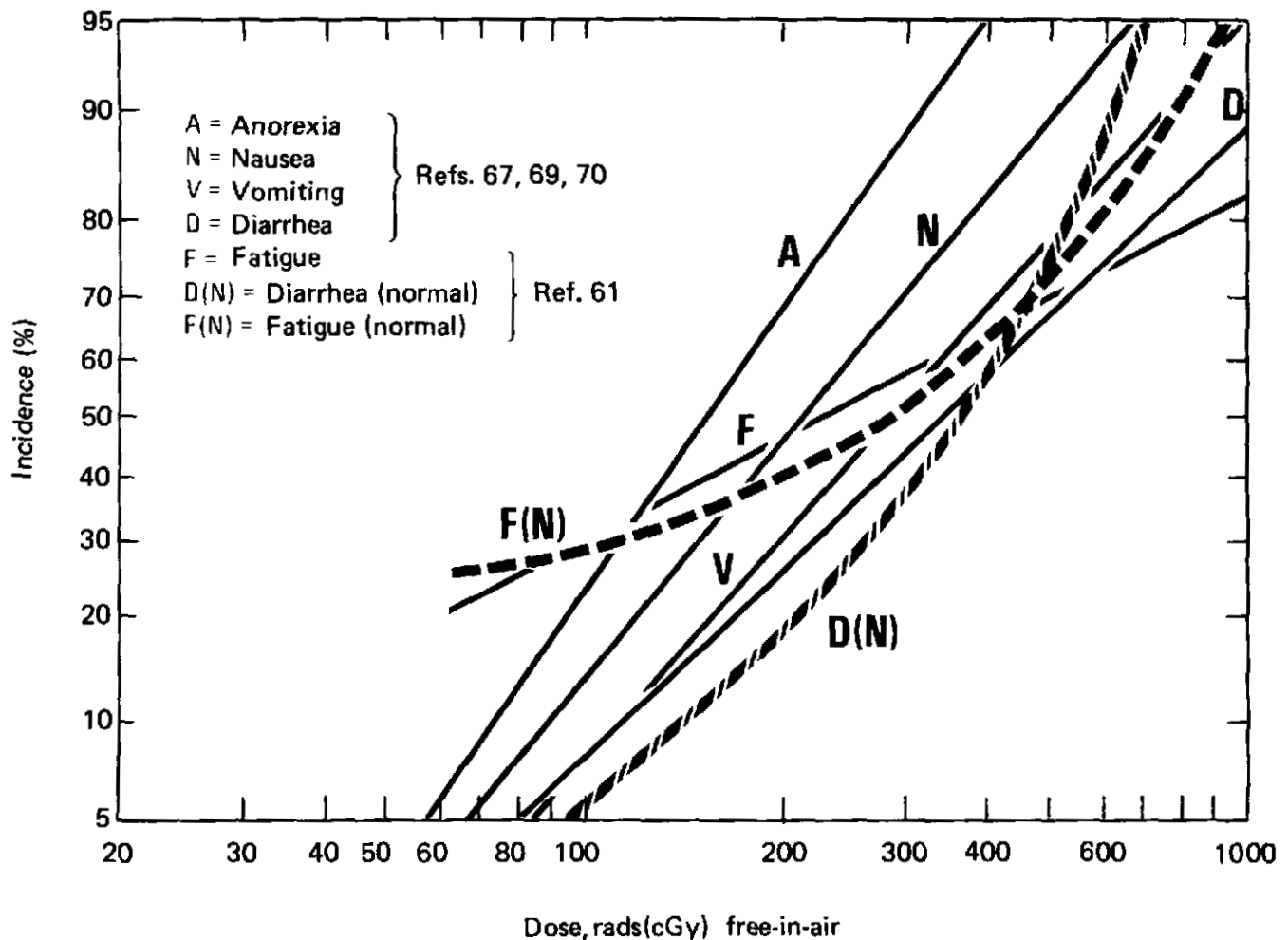


Figure 3. Symptom incidence related to dose (probit analyses).

fatigue at doses above 300 rads (cGy)* and the incidence of diarrhea at doses above 375 rads (cGy), for which the normal curves were used.

Qualification of the diarrhea data is in order. Unlike the other symptoms, diarrhea was monitored over a six-week period. Since the data are not time-resolved, the curves include later as well as early occurrences of diarrhea. In fact, other data suggest that these curves primarily reflect delayed diarrhea occurring a few days to a week after exposure. Early diarrhea is probably not manifested unless the dose is at least 450 rads (cGy). At that dose level, about 10 percent would

* Reference 61, Table 10, p. 82.

experience one or two episodes of diarrhea 3 to 6 hr after exposure; at doses of 3000 to 4500 rads (cgy), 30 percent could be affected.

RADIATION LETHALITY

Quantification of the incidence and postexposure time of death is uncertain because of the sparseness of human data. The three curves in Fig. 4 illustrate the uncertainty with respect to incidence: $LD_{50/60}$ ranges from 400 to 525 rads (cgy). The right-hand curve was obtained by matching an estimate of 525 rads (cgy) for humans with an $LD_{50/60}$ curve obtained from experiments with dogs. The right-hand curve was used because lethality-dose curves tend to be parallel for large animals.

For time of death after radiation exposure, the ranges indicated by the heavily outlined boxes in Fig. 5 were used. As the various grids show, the boxes represent a consensus of fairly recent expert opinion

and three fairly recent data points from human fatalities. For comparison a curve drawn through three earlier human data points is shown; the curve's shape is inferred from mammalian data. Relative to the curve, the boxes and the literature that support them show a trend toward earlier death with doses above ~2000 rads (cgy). That trend is also

consistent with the fatality reported at 35 hr after a dose of about 7000 rads (cgy).[†] The plateau in the curve may misrepresent human le-

thality because of differences between the gastrointestinal systems of man and other mammals. The hemopoietic, gastrointestinal, and central nervous system notations at the bottom of the figure indicate the ranges at which those syndromes are the major contributing causes of death.

SYMPTOM DESCRIPTIONS

Descriptions of the symptoms likely to be observed in humans after

exposure to ionizing radiation are presented for each of the eight dose ranges identified earlier. These descriptions, portrayed graphically

and summarized in text, estimate symptom onset, severity, duration, and

*
Reference 31.

†
References 38, 59.

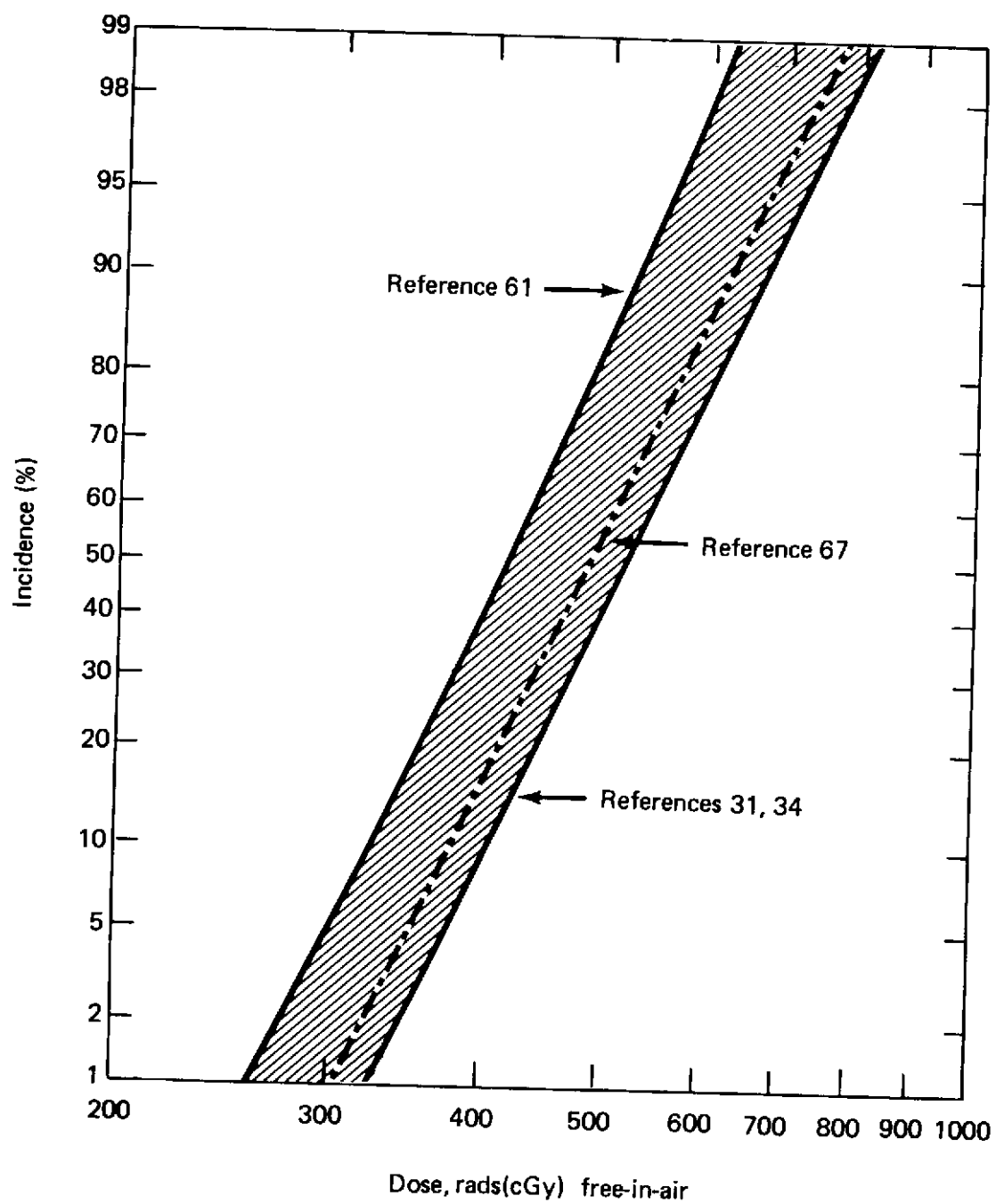
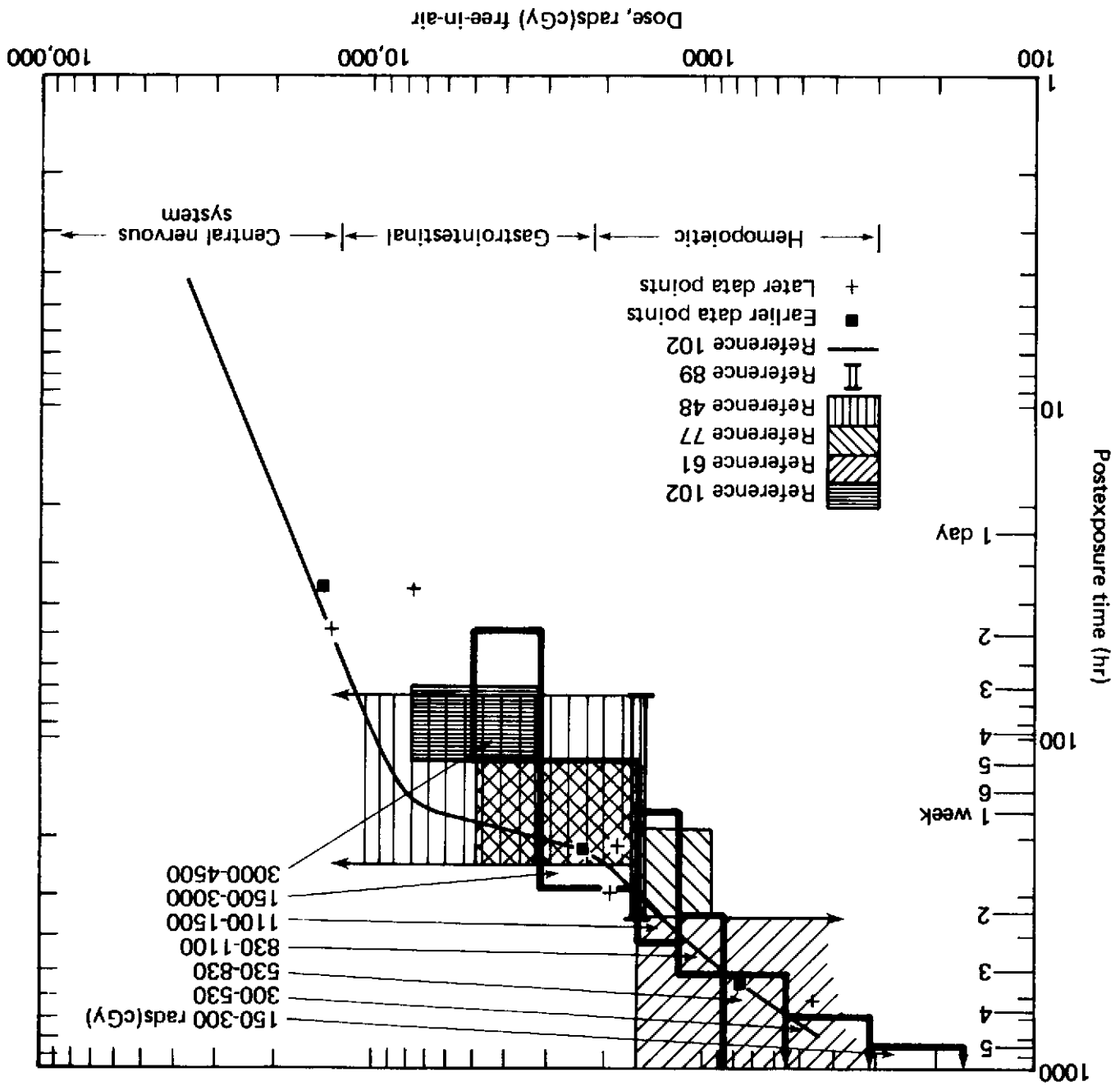


Figure 4. Lethality related to dose.

Figure 5. Time of death related to dose.



incidence. It should be reemphasized that a consensus was sought in the data, suppressing variability in order to represent the response of a typical exposed population.

Symptom incidence is expressed as a percentage range or single figure, depending on the firmness of the data. Estimates of the relation of symptom incidence with time are broad because of the lack of specific time-resolved data.

The estimates of duration imply that a symptom can occur one of three ways over the period indicated: continuously at one level of severity (e.g., anorexia); continuously with varying severity (e.g., fatigue^{*}); and intermittently (e.g., vomiting, diarrhea).

Quantification of the terms "mild," "moderate," and "severe" indicating symptom intensity is possible with only a few symptoms, as noted in Sec. 2.[†] Mild vomiting or diarrhea may mean a single to a few episodes during the period; moderate, several episodes; and severe, many and profuse episodes. Fatigue and weakness are potentially quantifiable since exertion, necessary to reveal those symptoms, is measurable;[‡] however, few data have been collected. With regard to hypotension, "mild" refers to a 10 percent drop in diastolic and systolic blood pressure; "moderate," a 10 to 30 percent drop; and "severe," a drop of 30 percent or more. Hemopoietic injury is quantified to the extent of estimating drops in platelet, granulocyte, and lymphocyte counts. Net continued fluid losses of up to 2 liters are considered mild to moderate; more than 2 liters, severe.

Regarding postexposure mortality, we estimate the incidence of fatalities and the period over which they are likely to occur for a given dose level. Data are insufficient to define a time distribution of mortality.

Doses of 75 to 150 Rads (cGy)

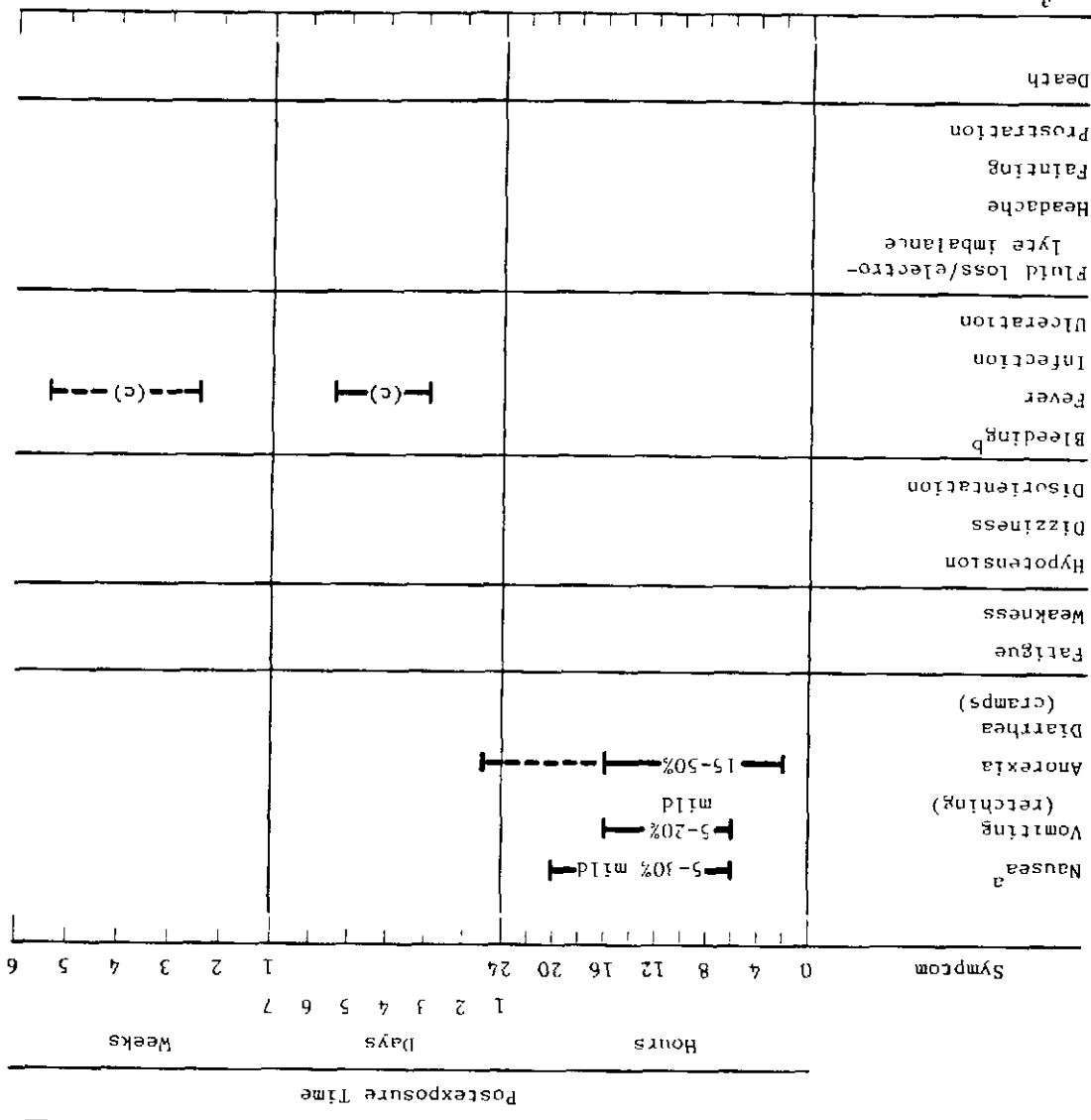
Table 2 indicates that acute radiation effects at this level are

^{*}Reference 21.

[†]Also, our correlations of severity level with time are gross representations; severity may vary considerably over time, as Gerstner's time-intensity profiles suggest (Refs. 41-43).

[‡]Reference 103.

Table 2. Symptoms for dose range 75 to 150 rads (cgy) free-in-air.



^aReferences for this group of symptoms: 1, 7, 15, 22, 26-30, 33, 37, 42, 50, 62, 66, 71, 72, 76, 78, 80, 85, 92, 96, 97, 105-107, 110, 111. These symptoms not observed in American servicemen exposed to approximately 78 rads (cgy) of fallout radiation, according to Reiss, 26-29.

^bReferences for this group of symptoms: 1, 7, 14, 15, 26-30, 33, 50, 64, 72, 101, 105, 106, 108.

^cSlight drop in lymphocyte, platelet, and granulocyte counts; no overt symptoms.

mild and occur only during the first day after exposure. Blood cell counts may drop slightly, but typical victims will surely survive.

Doses of 150 to 300 Rads (cGy)

The severity of prodromal effects increases over the preceding range (Table 3). As the dose approaches 200 rads (cGy), 50 percent or more of exposed persons develop anorexia, nausea, and vomiting. About 30 to 60 percent complain of fatigue and weakness. Significant destruction of bone marrow stem cells may lead to a 25 to 35 percent drop in blood cell production. As a result, mild bleeding, fever, and infection may occur during the fourth and fifth postexposure weeks. Up to 5 percent may die 5 to 6 weeks after exposure to 300 rads (cGy).

Doses of 300 to 530 Rads (cGy)

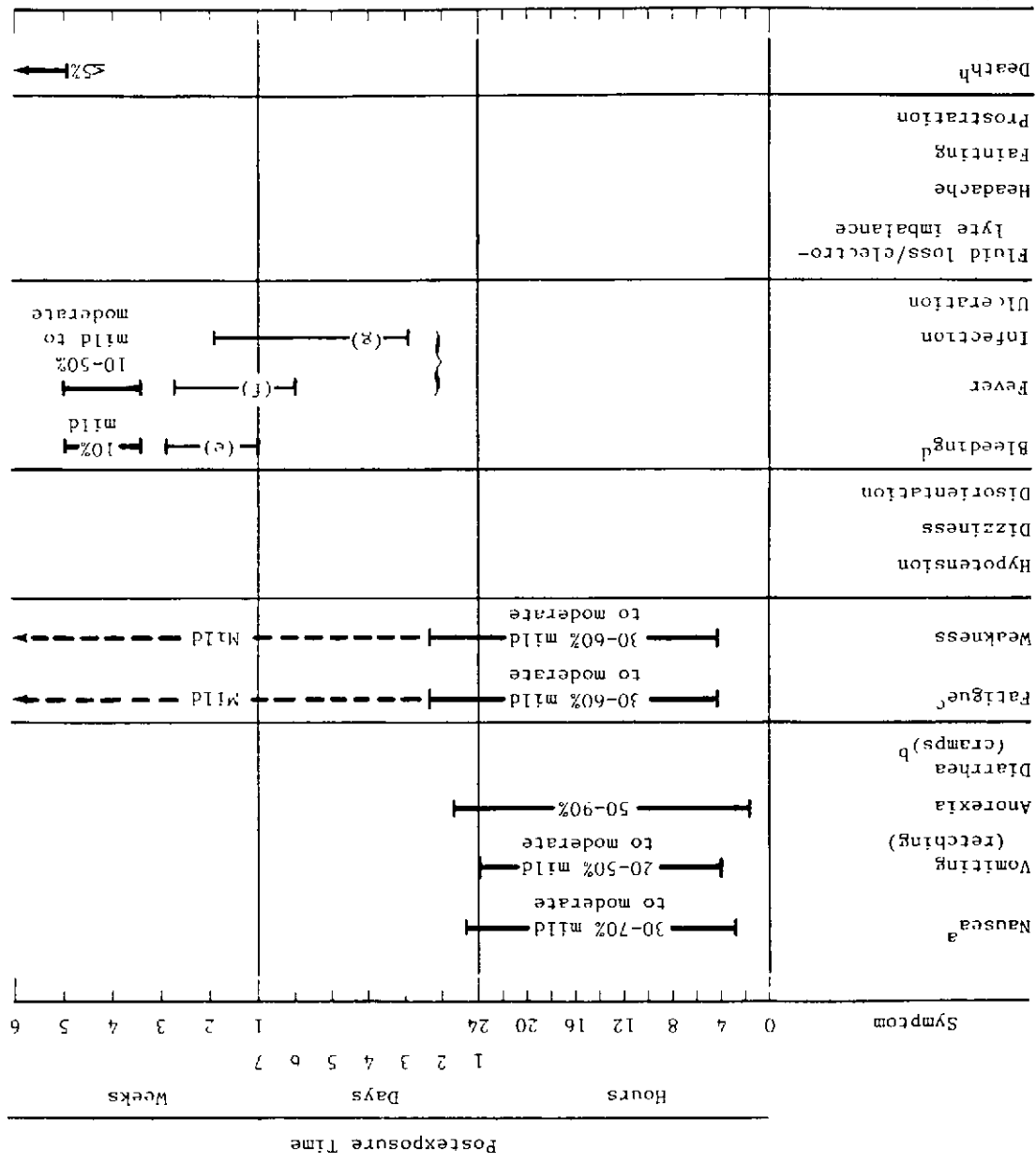
Prodromal symptoms begin earlier and affect more exposed persons (Table 4). At this range and above, 10 percent may experience one or two episodes of moderate diarrhea 4 to 6 hr postexposure. Most victims tire easily and experience mild to moderate weakness intermittently over the 6 weeks. Under normal conditions, vomiting and diarrhea are not enough to cause serious fluid loss and electrolyte imbalance. In hot or humid conditions, however, combined fluid loss and electrolyte imbalance could become serious.

Injury to the hemopoietic system is indicated by moderate bleeding, fever, infection, and ulceration 3 to 5 weeks postexposure; more than 50 percent of those exposed are affected. During the fourth and fifth weeks, moderate diarrhea may complicate their condition. Five to 50 percent of nontreated persons may die during the fifth week; death comes earlier to those with preexisting infections, for example of the upper respiratory tract.

Doses of 530 to 830 Rads (cGy)

The onset and duration of nausea, vomiting, and anorexia are about the same as in the preceding dose range, but the symptoms are more severe and affect nearly all exposed persons (Table 5). Severe and prolonged vomiting takes a toll on electrolyte balance, which would be

Table 3. Symptoms for dose range 150 to 300 rads (cGy) free-in-air.



^aReferences for this group of symptoms: 4, 6, 7, 15, 17, 21, 22, 26-29, 33, 37, 42, 44, 45, 50, 51, 56, 62, 65, 66, 74, 76, 78-82, 85-87, 90-92, 95, 96, 104, 105, 107, 111.

^bTen percent of the Marshallese victims exposed to 175 rads (cGy) experienced diarrhea during the first postexposure day, according to Ref. 4.

^cReferences for this group of symptoms: 7, 50, 60, 65, 81, 85, 86, 90, 101, 102.

^dReferences for this group of symptoms: 6, 7, 14, 18, 21, 25, 26-29, 33, 35, 60, 62, 64, 65, 75, 76, 78, 79, 81, 82, 85, 89, 101, 104, 105, 107, 110.

^eSlight to moderate drop in platelets: from $3 \times 10^5/\text{mm}^3$ to $1.8-0.8 \times 10^5/\text{mm}^3$.

^fSlight to moderate drop in granulocytes: from $6 \times 10^3/\text{mm}^3$ to $4.5-2.0 \times 10^3/\text{mm}^3$.

^gSlight to moderate drop in lymphocytes: from $3 \times 10^3/\text{mm}^3$ to $2.0-1.0 \times 10^3/\text{mm}^3$.

^hReferences for this event: 4, 14, 61, 81.

Table 4. Symptoms for dose range 300 to 530 rads (cGy) free-in-air.

| Symptom | Postexposure Time | | | | | | | | | | | | | | | | | | | |
|---------------------------------------|-------------------|---|---|----|----|----|----|---|------|---|---|---|---|---|---|-------|---|---|---|---|
| | Hours | | | | | | | | Days | | | | | | | Weeks | | | | |
| | 0 | 4 | 8 | 12 | 16 | 20 | 24 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 1 | 2 | 3 | 4 | 5 | 6 |
| Nausea ^a | | | | | | | | | | | | | | | | | | | | |
| Vomiting (retching) | | | | | | | | | | | | | | | | | | | | |
| Anorexia | | | | | | | | | | | | | | | | | | | | |
| Diarrhea (cramps) | | | | | | | | | | | | | | | | | | | | |
| Fatigue ^b | | | | | | | | | | | | | | | | | | | | |
| Weakness | | | | | | | | | | | | | | | | | | | | |
| Hypotension | | | | | | | | | | | | | | | | | | | | |
| Dizziness | | | | | | | | | | | | | | | | | | | | |
| Disorientation | | | | | | | | | | | | | | | | | | | | |
| Bleeding ^c | | | | | | | | | | | | | | | | | | | | |
| Fever | | | | | | | | | | | | | | | | | | | | |
| Infection | | | | | | | | | | | | | | | | | | | | |
| Ulceration | | | | | | | | | | | | | | | | | | | | |
| Fluid loss/electro- lyte imbalance | | | | | | | | | | | | | | | | | | | | |
| Headache | | | | | | | | | | | | | | | | | | | | |
| Fainting | | | | | | | | | | | | | | | | | | | | |
| Prostration | | | | | | | | | | | | | | | | | | | | |
| Death ^h | | | | | | | | | | | | | | | | | | | | |

^aReferences for this group of symptoms: 4, 6, 7, 12, 15, 18, 21, 22, 32, 33, 35, 37, 41-44, 47, 50, 51, 53, 56, 60, 62, 65, 75, 76, 80-82, 85-88, 90, 92, 95-97, 105-111.

^bReferences for this group of symptoms: 1, 7, 14, 43, 65, 71, 76, 81, 101, 104, 109, 110.

^cReferences for this group of symptoms: 2, 6, 7, 14, 15, 33, 35, 39, 54, 56, 60, 62, 64, 65, 71, 75, 76, 79, 81, 82, 85, 88, 95, 97, 101, 105, 106, 110.

^dModerate drop in platelets: from $3 \times 10^5/\text{mm}^3$ to $0.8-0.1 \times 10^5/\text{mm}^3$.

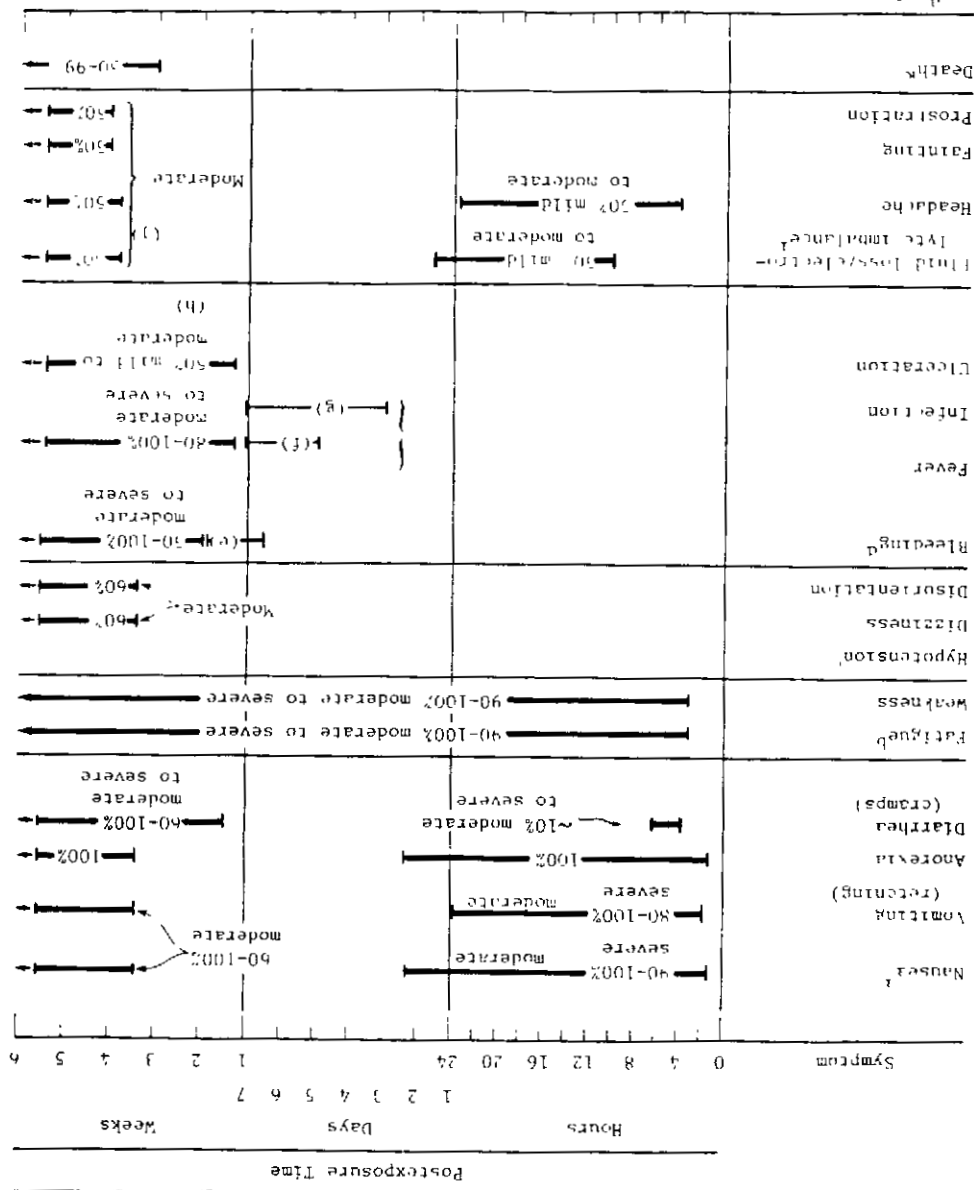
^eModerate drop in granulocytes: from $6 \times 10^3/\text{mm}^3$ to $2.0-0.5 \times 10^3/\text{mm}^3$.

^fModerate to severe drop in lymphocytes: from $3 \times 10^3/\text{mm}^3$ to $1.0-0.4 \times 10^3/\text{mm}^3$.

^gEpilation.

^hReferences for this event: 4, 7, 14, 61, 81.

Table 5. Symptoms for dose range 530 to 830 rads (cgy) free-in-air.



^aReferences for this group of symptoms: 1, 4, 6, 7, 14, 15, 21, 22, 32, 33, 37, 41, 42, 50, 51, 53, 56, 58, 62, 76, 77, 79-82, 85, 87, 89, 90, 95-97, 101, 103, 106, 110, 111.

^bReferences for this group of symptoms: 1, 6, 7, 14, 47, 51, 53, 65, 78, 81, 85, 90, 101.

^cReferences for this group of symptoms: 77, 89.

^dReferences for this group of symptoms: 1, 4, 7, 14, 15, 19, 31, 32, 33, 54, 58, 62, 64, 65, 67, 71, 75, 76, 77, 81, 85, 90, 95, 101, 103-107, 110, 111.

^eSevere drop in platelets: from $3 \times 10^3/\text{mm}^3$ to $0.1 \times 10^3/\text{mm}^3$.

^fSevere drop in granulocytes: from $6 \times 10^3/\text{mm}^3$ to $0.5 \times 10^3/\text{mm}^3$.

^gSevere drop in lymphocytes: from $3 \times 10^3/\text{mm}^3$ to $0.1 \times 10^3/\text{mm}^3$.

^hEpliation.

ⁱReferences for this group of symptoms: 7, 14, 21, 81, 89, 101.

^jMild intestinal damage.

^kReferences for this event: 4, 14, 61, 81.

accelerated by perspiration loss through heat, humidity, or activity. About 10 percent may experience moderate to severe diarrhea 3 to 6 hr after exposure. Nearly all show moderate to severe fatigue and weakness for many weeks. If untreated, 50 to 99 percent may die, primarily because of extensive injury to the hemopoietic system, manifested in overwhelming infections and bleeding during the third to sixth weeks. Nausea, vomiting, and anorexia may recur at that time. Diarrhea, electrolyte imbalance, and headaches affect at least half. The condition of the lethally irradiated during their last days may be complicated by dizziness, disorientation, fainting, prostration, and symptoms of infection and bleeding.

Doses of 830 to 1100 Rads (cGy)

Virtually all exposed persons experience severe nausea and vomiting the first postexposure day, moderating over the next day or two (Table 6). During that time they also become dizzy and disoriented.

With the near-maximum destruction of bone marrow stem cells and absence of granulocytes, untreated persons lose their defense against infection. By the end of the first postexposure week, infection is rampant from endogenous bacteria that have escaped from the injured gastrointestinal tract.

The combination of hemopoietic damage and gastrointestinal lesions reduces the survival of all untreated persons to 2 to 3 weeks. During the entire time they suffer from severe fatigue and weakness. Toward the end of the first week, nausea, vomiting, and anorexia recur. Moderate to severe diarrhea may begin as early as the fourth day. Severe bleeding, headaches, hypotension, dehydration, electrolyte imbalance, and fainting complicate the condition of all during their last days.

Doses of 1100 to 1500 Rads (cGy)

The survival time for untreated persons diminishes to 2 to 2½ weeks (Table 7). Symptoms resemble those experienced at the preceding dose range, with the following notable differences:

Table 6. Symptoms for dose range 830 to 1100 rads (cgy) free-in-air.

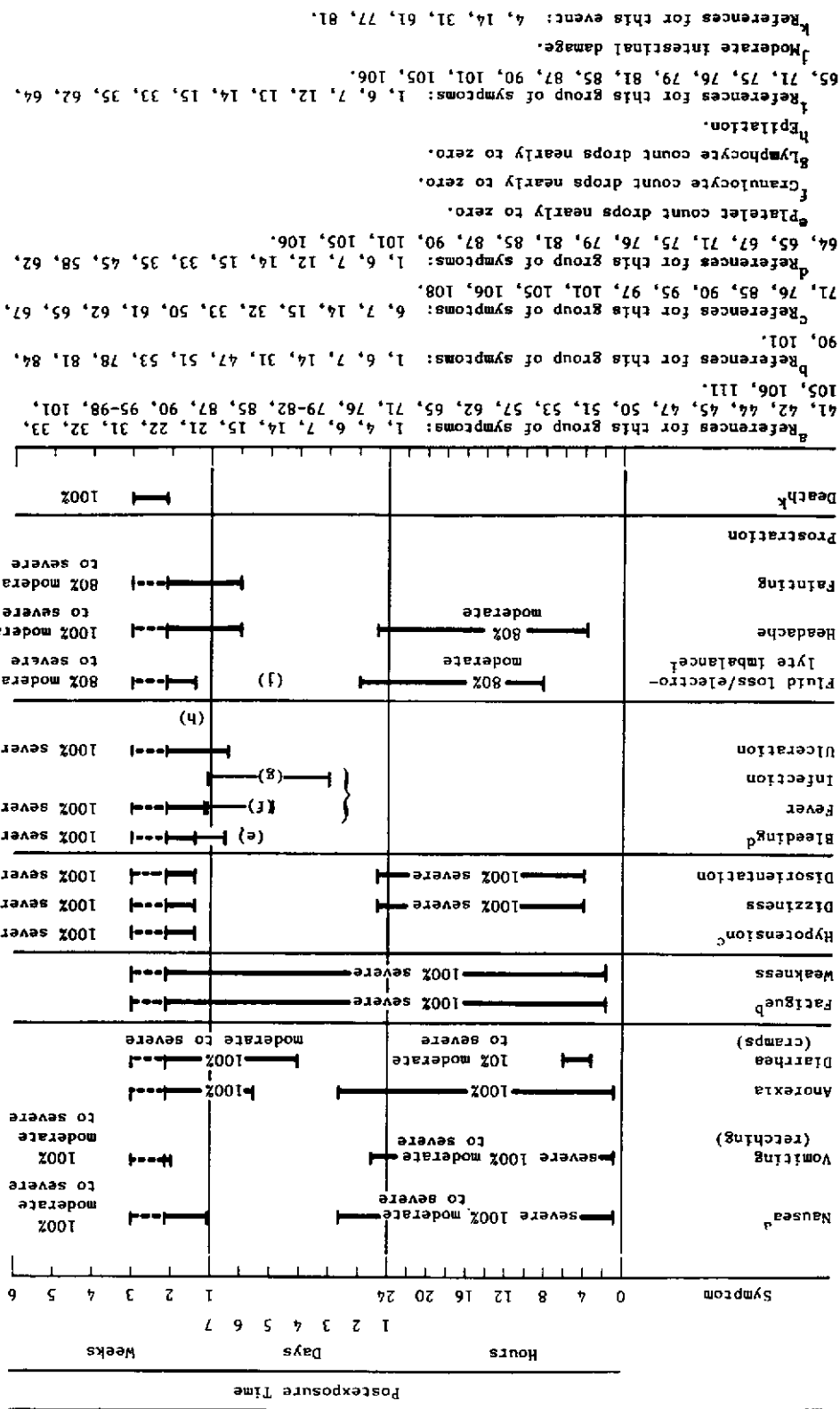
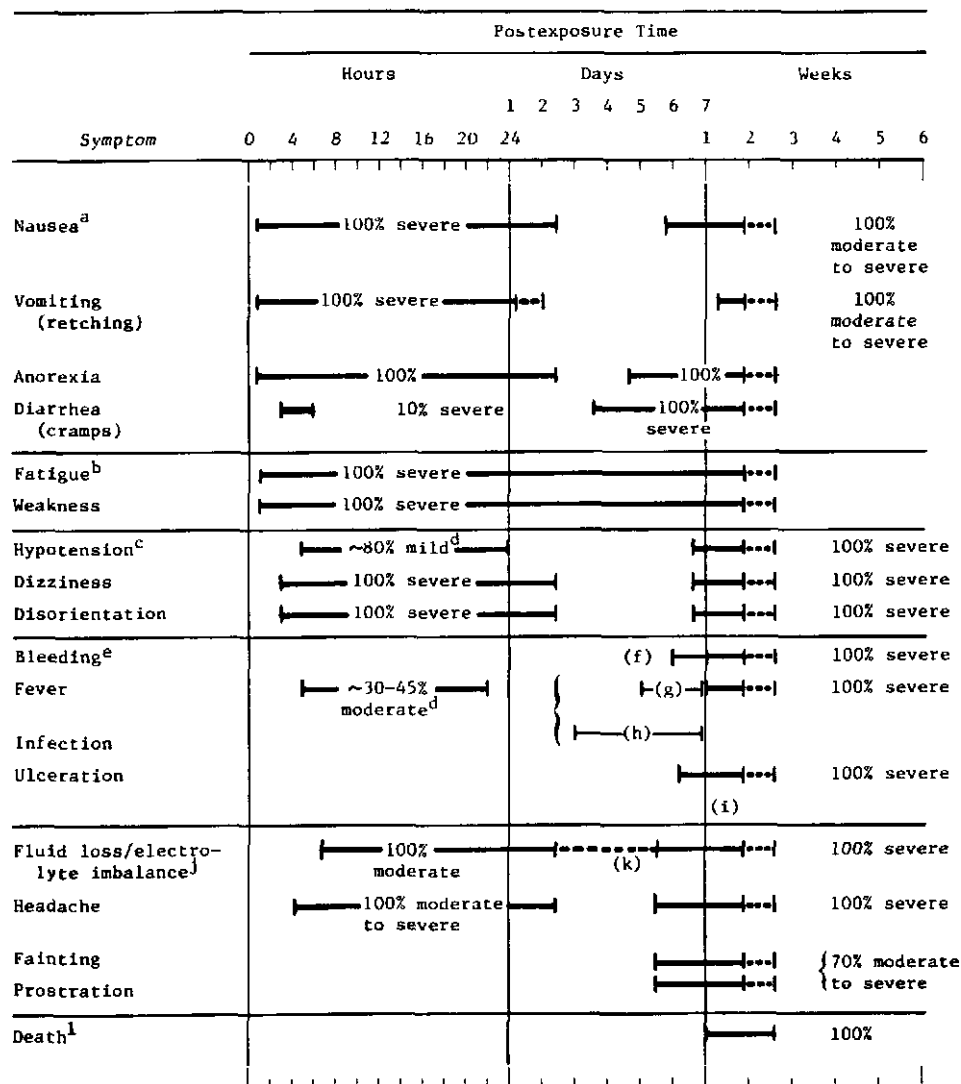


Table 7. Symptoms for dose range 1100 to 1500 rads (cGy) free-in-air.



^aReferences for this group of symptoms: 6, 7, 14, 15, 21, 22, 33, 41, 44, 47, 50, 51, 53, 57, 62, 65, 71, 76, 79-82, 85, 87, 90, 93, 95, 96, 101, 106.

^bReferences for this group of symptoms: 1, 6, 7, 14, 32, 47, 51, 53, 65, 67, 78, 85, 101.

^cReferences for this group of symptoms: 6, 7, 14, 15, 33, 50, 61, 62, 65, 71, 76, 79, 80, 82, 85, 87, 89, 93, 101, 105, 106.

^dBlood pressure drops 25 percent; temperature increases to 102°F, according to Ref. 33.

^eReferences for this group of symptoms: 6, 7, 14, 15, 21, 30, 31, 33, 51, 53, 61, 62, 65, 71, 72, 79-82, 85, 87, 89, 90, 93, 95, 101, 106.

^fPlatelet count drops to zero.

^gGranulocyte count drops to zero.

^hLymphocyte count drops to zero.

ⁱEpilation.

^jReferences for this group of symptoms: 6, 7, 14, 15, 33, 50, 61, 62, 65, 71, 72, 76, 79-82, 85, 89, 101, 106.

^kModerate to severe intestinal damage.

^lReferences for this event: 4, 14, 61, 77, 81.

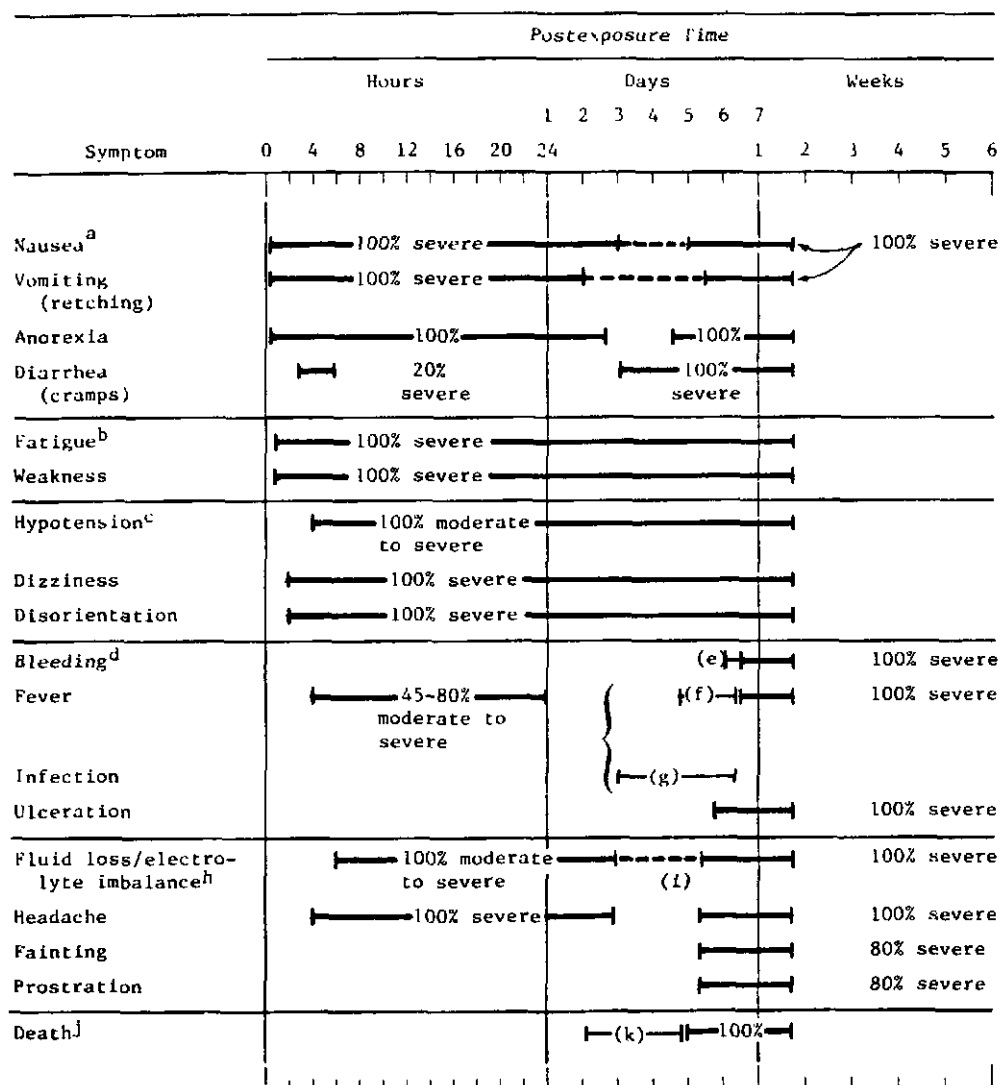
*Reference 85.
 †Reference 65. For more detail, see Appendix A.

Doses of 1500 to 3000 Rads (cgy)
 Severe nausea and vomiting affect all within 30 min of exposure and continue intermittently, along with anorexia, until death the second week (Table 8). Severe headaches begin after about 4 hr and continue for 2 to 3 days. Symptom severity may diminish somewhat during days 3 to 5. Gastrointestinal injury predominates, manifested 4 to 6 days after exposure by the abrupt return of severe nausea, vomiting, anorexia, and diarrhea, along with high fever, abdominal distension, and undetectable peristalsis (ileus).^{*} During the second week, severe dehydration, hemoconcentration, and circulatory collapse, compounded by septicemia, lead to coma and death.[†]

Much of the description of symptoms at this dose range derives from postexposure observations of patients treated with total-body irradiation for leukemia. There are undeniable difficulties in extrapolating from sick people under close medical attention to otherwise healthy young soldiers on the battlefield; those difficulties are discussed beginning on p. 35. Nevertheless, therapy patients constitute the only substantial number of irradiated persons whose reactions have been thoroughly documented, so their experience is relevant to this inquiry. The acute sequelae observed in therapy patients are detailed in Appendix B.

- Severe nausea and vomiting may continue into the third day before moderating.
- During the first day, hypotension affects about 80 percent; moderate fever, 30 to 45 percent.
- Electrolyte imbalance is a persistent problem from the sixth hour on.
- All have moderate to severe headaches during the first day.
- Nearly three-quarters are prostrate before the end of the first week.

Table 8. Symptoms for dose range 1500 to 3000 rads (cGy) free-in-air.



^aReferences for this group of symptoms: 6, 7, 14, 15, 21, 22, 31, 33, 47, 50, 62, 65, 67, 71, 76, 79-82, 85, 87, 90, 95, 101, 105, 106.

^bReferences for this group of symptoms: 6, 7, 14, 47, 53, 65, 71, 78, 85, 90, 101.

^cReferences for this group of symptoms: 6, 7, 14, 15, 33, 50, 61, 62, 65, 71, 76, 79-82, 85, 87, 89, 101, 105, 106.

^dReferences for this group of symptoms: 6, 7, 14, 15, 21, 22, 33, 47, 50, 62, 65, 67, 71, 76, 79-82, 85, 87, 90, 95, 101, 105, 106.

^ePlatelet count drops to zero.

^fGranulocyte count drops to zero.

^gLymphocyte count drops to zero.

^hReferences for this group of symptoms: 7, 14, 15, 33, 50, 61, 65, 68, 75, 76, 79-82, 85, 95, 101, 105, 106, 108.

ⁱSevere intestinal damage.

^jReferences for this event: 4, 14, 61, 81, 98.

^kRenal failure, according to Ref. 98.

Doses of 3000 to 4500 Rads (cgy)

Symptoms are more severe versions of those described for the preceding dose range (Table 9). Gastrointestinal injury predominates, complicated by cardiovascular lesions. Prodromal effects, including severe headache and drowsiness, appear almost immediately after exposure and may persist as the gastrointestinal syndrome develops. Severe dehydration and electrolyte imbalance are manifested several hours after exposure: initially fluids and electrolytes are lost by vomiting, but in time the greater loss is from severe diarrhea. The increased permeability of capillaries in the intestines and elsewhere in the body releases fluids into the interstitial spaces.*

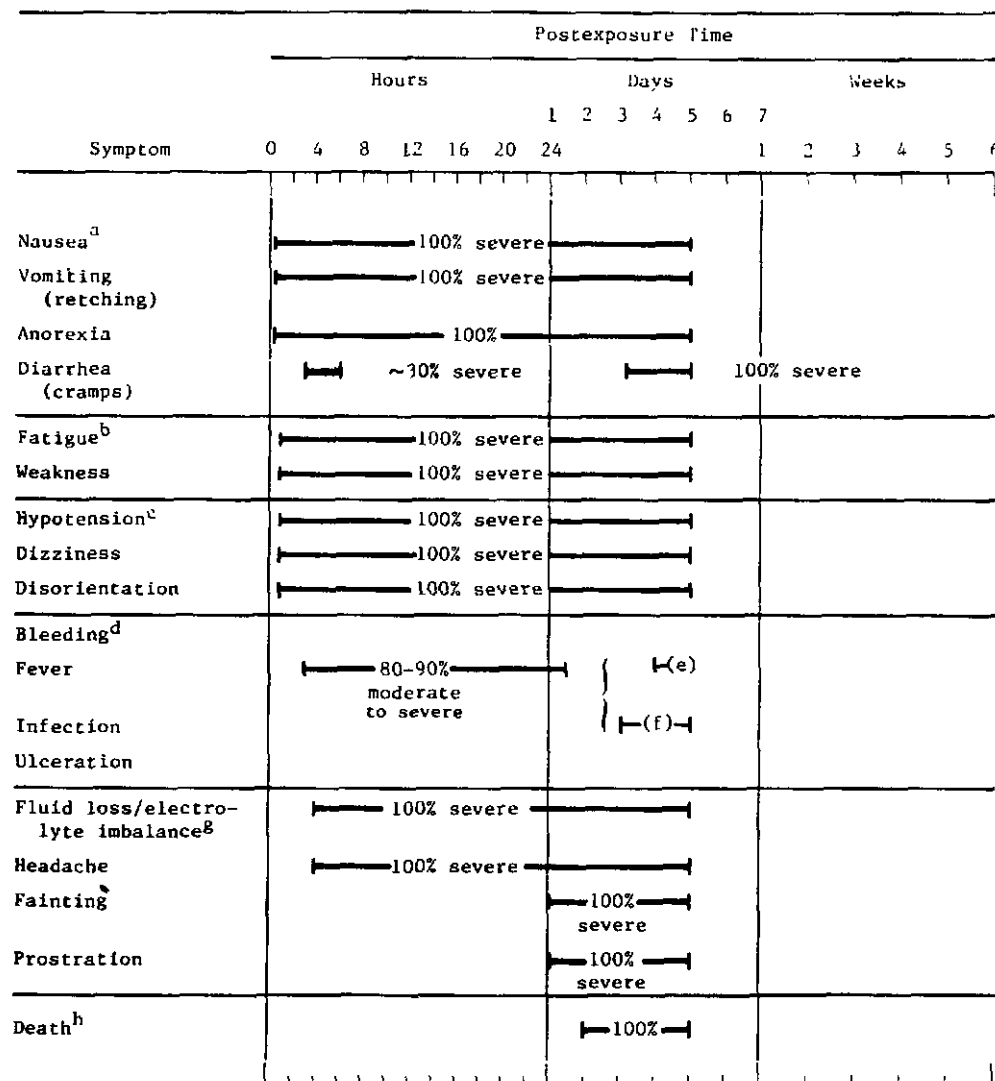
APPLICABILITY OF TYPICAL SYMPTOM DESCRIPTIONS

Application of the estimates in Tables 2 through 9 to battlefield soldiers raises several questions because of the obvious differences in population characteristics, environmental conditions, and medical attention. One question is whether the postexposure symptoms recorded for therapy patients could be distorted by the effects of their underlying disease and prior chemotherapy.

Another question pertains to the effects of postexposure medical care. Young soldiers trained for combat can generally be considered more robust than accident victims are before radiation exposure, and much more healthy than patients undergoing radiation therapy. Therefore, one might expect combat soldiers to better withstand the physical effects of ionizing radiation. However, all persons whose radiation response was examined in the data sources benefited from postexposure medical care, to which combat soldiers may not have access. Without such interventions as antibiotics and steroids, antiemetics, blood transfusions, intravenous fluids, bone marrow transplants, and bed rest, radiation sickness symptoms will be more widespread and severe, regardless of the individual's condition. The stresses engendered by nuclear war would of course compound the effects. Therefore, to what extent would the lack of medical care like that afforded

*References 68, 70, 85. For more detail, see Appendix A.

Table 9. Symptoms for dose range 3000 to 4500 rads (cGy) free-in-air.



^aReferences for this group of symptoms: 6, 7, 14, 15, 21, 22, 42, 44, 50, 62, 65, 71, 76, 79-82, 85, 87, 90, 95, 101, 106, 108.

^bReferences for this group of symptoms: 6, 7, 14, 15, 21, 22, 50, 62, 65, 71, 76, 79-82, 85, 87, 90, 94, 95, 101, 105, 106, 108.

^cReferences for this group of symptoms: 14, 50, 61, 71, 76, 79, 80, 82, 85, 87, 94, 95, 101, 105, 108.

^dReferences for this group of symptoms: 7, 14, 50, 61, 71, 75, 76, 79-82, 85, 87, 94, 95, 101, 105, 106, 108.

^eGranulocyte count drops to zero.

^fLymphocyte count drops to zero.

^gReferences for this group of symptoms: 14, 15, 33, 50, 61, 62, 71, 75, 76, 79, 80, 82, 85, 87, 95, 101, 105, 108.

^hReferences for this event: 4, 14, 61, 77, 81.

The ability of medical care to counter later somatic damage is demonstrated by the effectiveness of current bone marrow transplantation procedures in preventing death from bone marrow suppression after radiation therapy.

*

Given the absence of relevant quantitative data, definitive answers are as yet impossible. Let us then consider the matter qualitatively. Some have suggested that prodromal nausea, vomiting, fatigability, and diarrhea are partly psychogenic, so medical attention may be less effective in controlling them than in controlling purely somatic symptoms of hemopoietic or gastrointestinal damage. * Comparison of the experience of accident victims and therapy patients supports that assertion. When therapy patients received the best available antiemetic drugs before radiation, they showed similar patterns of early nausea and vomiting (incidence, severity, and duration) as accident victims exposed to comparable doses—even though the accident victims, of course, did not receive preirradiation antiemetics. The more recent introduction of steroid premedication in total-body radiation therapy markedly reduces the severity of prodromal effects. Of particular relevance in contrasting hospital care with battlefield situations is the reduced availability of intravenous fluid replacement in the field. Without that replacement, fluid loss and electrolyte imbalance can be a life-threatening result of severe vomiting (with or without diarrhea) or sweating from exertion, particularly in hot, humid conditions. For example, heat stroke is much more likely if a person is dehydrated from vomiting. Our general opinion is that preexposure health condition is less important than postexposure medical attention, barring prior bacterial or viral infection. Most experts would argue that, as dose increases beyond 300 rads (cgy) or so, any advantage gained through robustness will begin to be offset by the lack of medical care. As time increases past the first postexposure day, any such advantage would also diminish, given sublethal doses.

accident victims and therapy patients be offset by the prior youth,

vigor, and motivation expected in trained combat personnel?

With regard to dose, our symptom descriptions (based largely on the experience of persons who received medical care) are somewhat less applicable but still reasonably relevant to combat personnel exposed to doses at the lower end of our 75 to 4500 rad (cGy) range. The descriptions become increasingly applicable as dose increases.

With regard to postexposure time, prodromal symptoms are similar in both therapy patients (before steroids) and unpremedicated accident victims. The descriptions of manifest-illness symptoms draw mainly from the experience of accident victims and atomic bomb survivors. Therefore, the authors believe the estimates of symptom onset and duration given here apply reasonably well to battlefield personnel.

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PATHOPHYSIOLOGY OF RADIATION INJURY

Appendix A

This appendix supports and explains the findings reported in Sec. 3 by describing the pathophysiology of radiation injury. The acute period of injury is conventionally divided into prodromal and manifest-illness phases.

PRODROMAL PHASE

* The initial or prodromal phase begins about 2 to 4 hr postexposure for doses of 300 to 530 rads (cGy). As dose increases, the phase begins progressively earlier, within minutes of exposure to 4500 rads (cGy). Stomach distress appears, followed by anorexia. Vomiting occurs, especially with doses above 300 rads (cGy). If vomiting is severe, it is often accompanied by extreme weakness, and loss of body fluids can present problems. Depending on the dose, nausea, vomiting, and anorexia continue for 8 to 16 hr, even up to 2 days. Psychogenic factors may complicate the picture.[†] At lower dose ranges, a feeling of well-being may arise 24 to 48 hr after exposure and last 1 to 2 weeks.[‡]

Basic pathophysiological mechanisms in the prodromal phase remain somewhat unclear. Several causal factors have been suggested, including direct radiation effects on the central and autonomic nervous systems, disturbance of the endocrine balance, and production of various toxic substances.^{**} Symptoms of listlessness and nausea imply that higher structures of the nervous system are involved. However, since adult nervous systems are radioresistant in our dose range of interest, indirect involvement through chemical mediators is suspected.^{††}

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- * References 6, 7, 22, 47, 67, 81, 85, 89.
 - † References 7, 22, 47, 81.
 - ‡ Reference 85.
 - ** Reference 44.
 - †† Reference 44.

Vomiting is a complicated reflex between the central and autonomic nervous systems. Integrated in the medulla oblongata, the reflex is initiated by chemicals transported in the blood, impulses carried by autonomic fibers from abdominal organs to the medullary center, and the higher brain structures, including the cortex.^{*} Experiments by Chin and Wang identified the medulla's chemoreceptor trigger zone as the main source of postradiation nausea and vomiting.[†]

Thus, the prodromal gastrointestinal reaction, at least for doses up to 4500 rads (cGy), seems to result not from direct radiation effects on the nervous system but from chemical compounds acting mainly on the medullary trigger zone.[‡] It is thought that those chemical compounds are released from damaged cells of primarily lymphoid and bone marrow tissues.^{**} The cellular debris reaches a maximum at 8 to 12 hr post-exposure, and is cleared by fixed and free macrophages 24 to 48 hr post-exposure. It has been suggested that disintegration of the lymphoid tissue is a cause of all prodromal reactions.^{††} If so, phagocytic action may be largely responsible for limiting the duration of the prodromal phase for doses up to about 1500 rads (cGy). At doses of 1500 rads (cGy) and above, the metabolic breakdown of cells, proteins, and amino acids can release uric acid. Without the administration of intravenous fluids to maintain the proper pH level, uric acid crystals can precipitate in the urinary system.^{‡‡}

MANIFEST-ILLNESS PHASE

The later or manifest-illness phase is dominated by the hemopoietic and gastrointestinal syndromes. Much higher doses are required to initiate the gastrointestinal syndrome than the hemopoietic syndrome, but when both are present the compounding effect can be severe.

*Reference 16.

†Reference 24.

‡References 24, 44.

**References 36, 44.

††Reference 44.

‡‡Reference 8.

Hemopoietic Syndrome

At doses below 1050 rads (cgy), the hemopoietic syndrome begins about 8 to 10 days postexposure with a serious drop in granulocyte and platelet counts. * Pancytopenia supervenes about 3 to 4 weeks later; it becomes complete at doses above 750 rads (cgy). † Purpura is evident, and bleeding may be uncontrolled, causing anemia. Fever, pulse rate, and respiratory rate rise, due to endogenous bacterial and mycotic infections. ‡ Infections become uncontrolled given the impaired granulocyte and antibody production. † If at least 10 percent of the bone marrow stem cells remain uninjured, recovery is possible; otherwise, death occurs within 4 to 6 weeks. **

The pathophysiology of the hemopoietic syndrome is fairly well understood. †† All functional blood cells derive from common stem cells in the bone marrow, which are extremely radiosensitive. Functional cells such as granulocytes and platelets have a lifespan of only a few days, ‡ and adult cells outside the stem cell compartment cannot renew themselves. Therefore, when immature stem cells are destroyed, blood cell production plummets or stops. The death of granulocytes and platelets permits bleeding and ulceration, accompanied by fever and fatal infections. ***

Gastrointestinal Syndrome

At doses above 1050 rads (cgy), injury to the gastrointestinal tract contributes increasingly to the severity of the manifest illness phase. Under normal conditions, the integrity of the intestinal mucosa prevents the substantial escape of bacteria into the bloodstream. The

* References 2, 4, 18, 35, 57, 64, 75, 76, 106.

† References 6, 7, 81.

‡ References 30, 66, 111.

** References 7, 65, 81, 85.

†† References 2, 4, 7, 11, 14, 18, 32, 33, 37, 62, 65, 75, 85, 95, 97.

‡‡ References 11, 14, 41.

*** References 1, 7, 81, 101.

few bacteria that do escape are soon inactivated by granulocytes or specific antibodies. The radiosensitive mucosal stem cells in the crypts have a rapid turnover rate,^{*} producing mature, nondividing, differentiated cells that migrate to form the functional mucosal lining. Having a lifetime of several days, those mature mucosal cells are progressively shed and not replaced when radiation kills the stem cells in the crypt. The result is breakdown of the mucosa and ulceration. As the mucosa breaks down, large amounts of bacteria can enter the bloodstream. They go unchallenged because of the curtailed production of granulocytes[†] in the wake of radiation-caused damage to the bone marrow, so fever and infections are a consequence.[‡]

Even below the lower dose threshold cited above, at doses of about 450 to 1200 rads (cGy), temporary injury to the tight junctions between epithelial cells of the mucosal lining can permit the discharge of large amounts of molecular pyrogenic bacterial endotoxins into the bloodstream.^{**} At doses of 1050 to 1500 rads (cGy), the epithelial lining is more extensively depleted, and death results from septicemia within 2 to 3 weeks.^{††}

With doses of about 1500 to 2250 rads (cGy), denudation of the mucosa, particularly in the small intestine, leads to septicemia from gram-negative bacteria entering the bloodstream.^{‡‡} Beginning at doses of about 1900 rads (cGy), the septicemia is complicated by dehydration and electrolyte imbalance,^{***} resulting from exudation through the extensively ulcerated intestinal mucosa. Nutrition is lost because of impaired intestinal absorption, and infections are uncontrolled because of the complete bone marrow aplasia and subsequent pancytopenia.^{†††}

*References 14, 85.

†References 7, 65, 107.

‡References 13, 85.

**Reference 16.

††References 42, 50, 65, 81.

‡‡References 14, 33, 85.

***References 31, 64, 85.

†††References 14, 85.

This condition develops over a few days. Symptoms may not become severe until about the third or fourth day, when injury to the bone marrow and gastrointestinal tract described above leads to septicemia, fluid and electrolyte loss, and gastrointestinal infection accompanied by fever. Cramping, abdominal pains, and diarrhea, which may become watery, become more frequent and severe over the next week,* followed by shock and death.

At doses of about 2250 rads (cGy) and above, severe dehydration and electrolyte imbalance are likely to cause death from shock even before septicemia develops.

*References 7, 33, 65, 82, 85, 86, 111.

ACUTE SEQUELAE OF RADIATION THERAPY

Appendix B

This appendix draws on clinical experience with radiation therapy patients over the course of a decade to summarize the side effects of total-body irradiation.* The patients were being treated for leukemia, aplastic anemia, and other diseases, and received doses equivalent to single high-dose-rate exposures of about 750 to 1000 rads (cGy).[†] That is a dose range about which there is a dearth of information in other data sources, including case studies of accident victims, atomic bomb survivors, and early radiation therapy patients.

SYMPTOMS

Shortly after exposure, most patients experienced nausea, emesis, chills, and fever. Those symptoms usually subsided within about 10 hr and disappeared within 24 hr except for nausea and anorexia, which might persist for days. Emesis was aggravated by movement and often occurred with little warning.

In the first few hours after exposure, some patients showed decreased blood pressure and increased pulse rate due to circulatory hypovolemia. There were reports of acute myocardial insufficiency and death in patients with a history of myocardial disease.

A painful mumps-like swelling of the parotid gland developed within a few hours of exposure. The pain usually subsided within 2 days, but the swelling sometimes persisted for several more days. Xerostomia (dry mouth) sometimes lasted a week or more. During that time the saliva was reduced in volume, was thicker, and felt ropy. A metallic taste might persist as long as the mouth remained dry. Reduced salivary secretion added to patients' disinterest in food.

*The text is taken from Ref. 9, Appendix B.

[†]Unless otherwise noted, doses in this appendix are absorbed mid-line tissue doses. To convert to free-in-air dose values, multiply by 1.5.

About 10 percent of the patients developed diarrhea soon after irradiation. More developed diarrhea 1 to 7 days after exposure. Fatigability during the first 24 hr might be related to fluid loss and electrolyte disturbance. Thereafter the patient might feel better temporarily, but usually after 3 or 4 days diarrhea developed. At the same time, a sense of easy fatigability, apparently unrelated to fluid and electrolyte changes, sometimes returned to become a major complaint.

The oropharyngeal mucosae became reddened and sore 1 to 3 days after exposure and subsequently ulcerated. The condition took about 3 weeks to disappear. About 75 percent of the patients developed oral infections, owing not only to the ulcerated mucosae but also to leukocytopenia and immunosuppression. These infections became apparent as soon as 3 days after irradiation. The most common were fungal (thrush), but bacterial and herpes infections were also seen. Bacterial infections would probably have been more common except that patients were given antibacterial drugs.

Bone marrow suppression was indicated by increased susceptibility to infections and bleeding (e.g., of the gums) several days after exposure. If the patient had a preexisting infection, however, total-body irradiation was usually fatal, sometimes during the first postexposure week. Bone marrow grafts did not help.

A generalized erythema appeared as soon as 1 day after irradiation though usually later. It persisted for as long as 2 weeks and was sometimes associated with perineal irritation and itchiness. Beginning 7 to 10 days after exposure there was a temporary incomplete loss of hair.

Sweating appeared to decrease in some patients. Though that phenomenon has not been adequately investigated, we can make a few observations. Inhibition of sweating seems not to present a serious problem in usual X- or gamma-ray treatment. The energy of the beam is high enough to spare the skin, and only small surface areas are irradiated. Inhibition of sweating could be lethal if large single doses are delivered from sources that do not spare the skin. Electrons were used to treat the total skin surface of a group of patients with mycosis fungoides. Penetrating only ~1 cm below the skin, the electron radiation was equivalent

to a single acute dose of 1000 to 2000 rads (cgy). Patients experienced marked erythema, decreased sweating associated with a generalized burn-sensation, and low tolerance to exercise with consequent risk of hyperthermia.

PROBLEMS IN ASSESSING SYMPTOMS

Advances in therapeutic methods have introduced multiple variables that complicate symptomatology assessment. Some side effects (e.g., oral mucosal) may be intensified by the prior administration of cytotoxic chemotherapy drugs. Acute side effects have been markedly alleviated by premedication with antiemetics, steroids, and intravenous fluids. Infections have been reduced by preirradiation decontamination and by not treating patients having evidence of infection.

All patients undergoing total-body irradiation received postirradiation bone marrow transplants and drugs to combat graft-versus-host disease (GVHD). Since the anti-GVHD drugs produce side effects similar to those induced by radiation, it is difficult to distinguish the radiation-unique effects.

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 ATTN: OSIB
 ATTN: RTS-2B
 ATTN: RTS-2C, Tech Svcs & Spt
 ATTN: DB-4, Rsch, Resources Div
 Defense Intel College
 ATTN: DIC/RTS-2
 ATTN: DIC-2C
 Defense Logistics Agency
 ATTN: Command Security Ofc
 Defense Nuclear Agency
 ATTN: CID
 ATTN: NASF
 ATTN: NATF
 ATTN: NAME
 ATTN: OASIS
 ATTN: RAE
 ATTN: STNA
 ATTN: STRA
 ATTN: STSP
 ATTN: OADP
 ATTN: STTI-CA
 4 cys ATTN:
 216 cys ATTN: OPNS
 National Security Agency
 ATTN: A12, F. Newton
 ATTN: Chief, A Group
 National Defense University
 ATTN: ICAF, Tech Lib
 ATTN: NMCLB-CR
 ATTN: Stop 315, Library
 ATTN: Strat Concepts Div Ctr
 Joint Data System Support Ctr
 ATTN: C-312
 ATTN: C-332
 ATTN: C-343
 Joint Strat Tgt Planning Staff
 ATTN: JLS
 ATTN: JLT
 ATTN: JP
 ATTN: JPPFD
 ATTN: JPTP
 2 cys ATTN: JLK, DNA Rep
 Joint Chiefs of Staff
 ATTN: ED30, J-3, Strat Ops Div
 ATTN: GD10, J-5, Nuc & Chem Div
 ATTN: J-3, Strat Ops Div
 ATTN: J-5, Strat Div, W. McClain
 ATTN: JAD/SFD
 ATTN: JAD/SSD
 ATTN: JSDA
 ATTN: SAGA
 ATTN: J-3, Special Opns
 ATTN: J-5, Nuc Div/Strat Div/FP&P Div
 ATTN: J-5, Plans & Policy/Nuc Chem Div
 Joint Data System Support Ctr
 ATTN: C-312
 ATTN: C-332
 ATTN: C-343
 Joint Strat Tgt Planning Staff
 ATTN: JLS
 ATTN: JLT
 ATTN: JP
 ATTN: JPPFD
 ATTN: JPTP
 2 cys ATTN: JLK, DNA Rep
 National Defense University
 ATTN: ICAF, Tech Lib
 ATTN: NMCLB-CR
 ATTN: Stop 315, Library
 ATTN: Strat Concepts Div Ctr
 National Security Agency
 ATTN: A12, F. Newton
 ATTN: Chief, A Group

DEPARTMENT OF DEFENSE (Continued)

Ofc of the Sec of Def, Net Assessments
ATTN: Doc Control

Program Analysis & Evaluation

ATTN: Naval Forces
2 cys ATTN: Regional Programs
2 cys ATTN: Strategic Programs

US European Command

ATTN: ECJ-LW
ATTN: ECJ-2
ATTN: ECJ-2-ITD
ATTN: ECJ-3
ATTN: ECJ-5
ATTN: ECJ5-N, Nuc Div
ATTN: ECJ-6
ATTN: ECJ2-T, Tgts Div
2 cys ATTN: ECCS/SASM
2 cys ATTN: ECJ-7 LW

US National Mil Representative, SHAPE

ATTN: US Documents Officer

Under Sec of Def for Policy

ATTN: DUSP/P
ATTN: USD/P

Under Secy of Def for Rsch & Engrg

ATTN: Tactical Warfare Prog
ATTN: Chairman, Def Sci Brd
ATTN: Strat & Space Sys (OS)
ATTN: Strat & Theater Nuc Forces, F. Vajda
2 cys ATTN: C3I
10 cys ATTN: Chairman, PSEAG

DEPARTMENT OF THE ARMY

Army Research Institute

ATTN: Commander

BMD Program Office

ATTN: DACS-BM, J. Kahlas, 13101

Combat Material Eval Element

ATTN: Security Analyst

Dep Ch of Staff for Ops & Plans

ATTN: DAMO-NC, Nuc Chem Dir
ATTN: DAMO-NCN
ATTN: DAMO-RQS
ATTN: DAMO-ZXA

Dep Ch of Staff for Rsch, Dev & Acq

ATTN: DAMA-CSS-N

Dept of the Army

ATTN: DAMA-CSS-N
ATTN: DAMI-CI
ATTN: DAMO-NCZ
ATTN: DAMO-OD
ATTN: DAMO-ODSO
3 cys ATTN: DAPE-HRE

Eighth US Army

ATTN: CJ-POX-NS

Harry Diamond Laboratories

ATTN: DELHD-NW-P
ATTN: DELHD-TA-L, 81100, Tech Lib

DEPARTMENT OF THE ARMY (Continued)

Joint Strategic Opns Ctr

2 cys ATTN: J-2
2 cys ATTN: J-5

Military Traffic Mgt Command

ATTN: Ofc of Security & Safety

Seneca Army Depot

ATTN: Provost Marshal
ATTN: SDSSE-PO
ATTN: Surety

Sierra Army Depot

ATTN: Security Opns

Southern European Task Force

ATTN: AESE-GCT-S

US Army Air Defense School

ATTN: Commandant

US Army Armament Rsch Dev & Cmd

2 cys ATTN: DRDAR-LCN-F

US Army Armor School

ATTN: ATSB-CTD
ATTN: Tech Library

US Army Ballistic Research Lab

ATTN: AMXBR-VLD, Dr. Klopac
ATTN: DRDAR-BL
ATTN: DRDAR-BLA-S, Tech Lib
ATTN: DRDAR-BLT
ATTN: DRDAR-BLV-R, Dr. Rainis

US Army Belvoir R&D Ctr

ATTN: DRCPM-PSE

US Army Comb Arms Combat Dev Acty

ATTN: ATZL-CAP

US Army Comd & General Staff College

ATTN: Acq Library Div
ATTN: ATSW-TA-D
ATTN: ATZL-SWJ-CA
ATTN: ATZL-SWS-L, D. Dorris

US Army Concepts Analysis Agency

ATTN: CSSA-ADL, Tech Lib

US Army Corps of Engineers

ATTN: Security & Law Enforcement

US Army Criminal Investigation Cmd

ATTN: Commander

US Army Elct Warfare Lab

ATTN: DELEW-I-S

US Army Electronic Proving Ground

ATTN: STEEP-PA-I

US Army Europe & Seventh Army

ATTN: AEACC-ND
ATTN: AEAGC
ATTN: Provost Marshal
2 cys ATTN: AEAGD-MM-SW
2 cys ATTN: AEAPM-PS
2 cys ATTN: DCSI
2 cys ATTN: DCSOPS

DEPARTMENT OF THE ARMY (Continued)

VII Corps
ATTN: G-3

1st Special Operations Command
ATTN: AFVS-GC-0, Maj Ogden

59th Ordnance Brigade
ATTN: AEUSA-Z
ATTN: Surety

DEPARTMENT OF THE NAVY

Carrier Airborne Early Warning Wing 12
ATTN: Commander

Carrier Group 1
ATTN: Commander

Carrier Group 2
ATTN: Commander

Carrier Group 3
ATTN: Commander

Carrier Group 4
ATTN: Commander

Carrier Group 5
ATTN: Commander

Carrier Group 6
ATTN: Commander

Carrier Group 7
ATTN: Commander

Carrier Group 8
ATTN: Commander

Cruiser-Destroyer Group One
ATTN: Commander

Cruiser-Destroyer Group 12
ATTN: Commander

Cruiser-Destroyer Group 2
ATTN: Commander

Cruiser-Destroyer Group 3
ATTN: Commander

Cruiser-Destroyer Group 5
ATTN: Commander

Cruiser-Destroyer Group 8
ATTN: Commander

David Taylor Naval Ship R&D Ctr
ATTN: Code 174

Fighter Airborne Early Warning Wing, US Pacific Fleet
ATTN: Commander

Fighter Wing 1
ATTN: Commander

Fleet Intelligence Center, Pacific
ATTN: FICPAC, Code 21

DEPARTMENT OF THE ARMY (Continued)

US Army Field Artillery School
ATTN: ATSF-CD

US Army Forces Command
ATTN: AF-OPTS

US Army Human Engineering Lab
ATTN: Director

US Army Infantry Ctr & Sch
ATTN: ATSH-CD-CSO

US Army Intel Threat Analysis Det
ATTN: IAX-Z

US Army Intelligence Agency
ATTN: DELEW-I

US Army Material Command
ATTN: DRCPM-NUC

US Army Material Command
ATTN: DRCD-DE-D

US Army Material Command
ATTN: DRCC

US Army Material Command
ATTN: DRXSY-DS

US Army Military Police School
ATTN: ATZN-MP-Library

US Army Military Police School
ATTN: ATZN-MP-TD

US Army Nuc & Chem Agency
ATTN: Library

US Army TRADOC Sys Analysis Acty
ATTN: ATAA-TAC

US Army Training & Doctrine Command
ATTN: ATCD-AO

US Army Training & Doctrine Command
ATTN: ATCD-FA

US Army Training & Doctrine Command
ATTN: ATCD-N, Cbt Dev, Nuc Dir

US Army Training & Doctrine Command
ATTN: ATCD-NCO

US Army War College
ATTN: Library

US Army War College
ATTN: Strategic Studies

US Army War College
ATTN: Strategic Studies

US Army War College
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US Army War College
ATTN: Strategic Studies

DEPARTMENT OF THE NAVY (Continued)

Fleet Intelligence Ctr, Europe & Atlantic
ATTN: Library

Naval Material Command
ATTN: MAT 0433
ATTN: MAT-0462
ATTN: PM-23

Light Attack Wing, US Pacific Fleet
ATTN: Commander

Light Attack Wing 1
ATTN: Commander

Marine Corps
ATTN: Code PPO

Marine Corps Dev & Education Command
ATTN: Commander

Medium Attack Tactical Electronic Warfare Wing
US Pacific Fleet
ATTN: Commander

Medium Attack Wing 1
ATTN: Commander

Naval Air Force, US Atlantic Fleet
ATTN: Commander

Naval Air Force, US Pacific Fleet
ATTN: Commander

Naval Electronic Sys Engineering Center
ATTN: Code 04
ATTN: Code 404HS

Naval Facilities Engineering Command
ATTN: Code 032E

Naval Intelligence Support Ctr
ATTN: NISC-30

Naval Investigative Svcs
ATTN: NISHC-22A
ATTN: NOP-009D
ATTN: 009/NIS/243

Naval Ocean Systems Center
ATTN: Code 4471, Tech Lib

Naval Personnel Res & Dev Ctr
ATTN: Code P302

Naval Postgraduate School
ATTN: Code 1424, Library

Naval Research Laboratory
ATTN: Code 1240
ATTN: Code 2627, Tech Lib

Naval Sea Systems Command
ATTN: SEA-09G53, Lib
ATTN: SEA-643

Naval Surface Force, US Atlantic Fleet
ATTN: Commander

Naval Surface Force, US Pacific Fleet
ATTN: Commander

DEPARTMENT OF THE NAVY (Continued)

Naval Surface Weapons Center
ATTN: Code F31
ATTN: G, G00

Naval War College
ATTN: Code E-11, Tech Svc
ATTN: Ctr for Nav Warfare Studies
ATTN: Doc Control
ATTN: Library
ATTN: Strategy Dept

Naval Weapons Evaluation Facility
ATTN: Tech Director

Nuc Wpns Tng Group, Atlantic
ATTN: Code 222
ATTN: Doc Control

Nuclear Weapons Tng Group, Pacific
ATTN: Code 32
ATTN: Doc Control

Ofc of the Dep Ch of Naval Ops
ATTN: NIS-22
ATTN: NOP CC9D
ATTN: NOP CC9D3
ATTN: NOP C6D
ATTN: NOP 50, Avn Plns & Rqmts Dev
ATTN: NOP 60
ATTN: NOP 60D
ATTN: NOP 603
ATTN: NOP 654, Strat Eval & Analysis Br
ATTN: NOP 91
ATTN: NOP 955, AAW Div
ATTN: NOP 981
2 cys ATTN: NOP 403

Ofc of Naval Research
ATTN: Code 713

CNO Exec Panel, Ofc of the Ch of Naval Opns
ATTN: OP-00K

Operational Test & Eval Force
ATTN: Commander

Operational Test & Eval Force, Pacific
ATTN: Dep Commander

Dep Ch of Staff, Plans, Policy & Opns
ATTN: Code-P
ATTN: Code-PDC-30

Space & Naval Warfare Systems Cmd
ATTN: PME 121-3

Strategic Systems Programs, PM-1
ATTN: Code SP113

Submarine Force, US Atlantic Fleet
ATTN: Commander

Submarine Force, US Pacific Fleet
ATTN: Commander

Submarine Group 2
ATTN: Commander

Submarine Group 5
ATTN: Commander

DEPARTMENT OF THE NAVY (continued)

Submarine Group 6
ATTN: Commander

Submarine Group 7
ATTN: Commander

Submarine Group 8
ATTN: Commander

Submarine Group 9
ATTN: Commander

Tactical Ing Gp, Pacific
ATTN: Commander

Tactical Wings Atlantic
ATTN: Commander

Commander in Chief, US Atlantic Fleet
ATTN: J2
ATTN: Physical Security
ATTN: Plans & Operations

Commander in Chief, US Naval Forces, Europe
ATTN: NS4, Nuc Warfare Officer
ATTN: Special Ops

US Navy Second Fleet
ATTN: Commander

US Navy Seventh Fleet
ATTN: Commander

US Navy Sixth Fleet
ATTN: Commander

US Navy Third Fleet
ATTN: Commander

Commander in Chief, US Pacific Fleet
ATTN: J-2
ATTN: Physical Security
ATTN: Plans & Ops

DEPARTMENT OF THE AIR FORCE

Aeronautical Systems Division
ATTN: XRO/MAF

Air Force
ATTN: INA

Air Force
ATTN: INT

Air Force Logistics Command
ATTN: Security Police

Air Force Ofc of Special Investigations
ATTN: IVS

Air Force Office of Security Police
2 cys ATTN: AFOSP/SPPC
2 cys ATTN: AFOSP/SPPX
2 cys ATTN: SPOS-SPPC

Air Force Systems Command
ATTN: DL
ATTN: SD
ATTN: Security Police
ATTN: XR

DEPARTMENT OF THE AIR FORCE (continued)

Air Force Weapons Laboratory
ATTN: SUL

Air Training Command
ATTN: Security Police

Air University
ATTN: AU/SP
ATTN: Strategic Studies

Air University Library
ATTN: AUL-LSL
ATTN: Library

Assist Ch of Staff, Studies & Analysis
2 cys ATTN: AF/SAMI, Tech Info Div

Assist Ch of the Air Force, Resch, Dev & Logistics
ATTN: SAF/ALR

Dep Ch of Staff, Resch, Dev & Acq
ATTN: AF/RDQ1

Dep Ch of Staff, Plans & Ops
ATTN: AF/XOIR
ATTN: AF/XOIFM, Plans, Frc Dev Mun Plans
ATTN: AF/XOIFS, Frc Dev, Strat Off Frc

Electronic Systems Division
3 cys ATTN: Physical Security Sys Directorate

Foreign Technology Division
ATTN: CCN
ATTN: SDN
ATTN: TQTM

Military Air Lift Command
ATTN: Security Police

Commander in Chief, Pacific Air Forces
ATTN: Security Police
ATTN: XP

Space Command
ATTN: Security Police

Space Division
ATTN: YH, DSCS III

Strategic Air Command
ATTN: Security Police
ATTN: ADMN
ATTN: NRI/STINFO
ATTN: SPD
ATTN: STIC, 544SIW
ATTN: XOXO
ATTN: XPQ
ATTN: XPZ

Tactical Air Command
ATTN: Security Police
ATTN: TAC/XPJ
ATTN: TAC/XPS

US Air Force Academy
ATTN: Library
ATTN: Strategic Studies
ATTN: USAFA/SP

US Air Force in Europe
2 cys ATTN: USAFE/SP

DEPARTMENT OF THE AIR FORCE (Continued)

US Air Force Inspector General

3 cys ATTN: IGS

3 cys ATTN: IGT

US Air Forces in Europe

ATTN: USAFE/DEX

ATTN: USAFE/DOT

ATTN: USAFE/INAT

ATTN: USAFE/XPX, Plns

USAF School of Aerospace Medicine

ATTN: Radiation Sciences Div

USAF Special Operations School

ATTN: Director

1st ACCS

ATTN: DOF

2nd ACCS

ATTN: Doc

3280th Tech Training Sq

ATTN: TG1CC

DEPARTMENT OF ENERGY

Department of Energy

Albuquerque Operations Office

ATTN: CTID

ATTN: D. Richmond

Department of Energy

Office of Mil Application, GTN

ATTN: OMA, DP-22

Department of Energy, GTN

ATTN: Ofc of Intelligence

ATTN: OMA, DP-22

ATTN: Safeguards & Security

ATTN: Tech & Intell Dir

University of California, Lawrence Livermore Natl Lab

ATTN: L-35

ATTN: L-38

ATTN: L-389

ATTN: L-450, W. Hogan

ATTN: Tech Info Dept Lib

ATTN: Z Division Library

Los Alamos National Laboratory

ATTN: M/S634, T. Dowler

ATTN: MS P364, Reports Library

ATTN: R. Sandoval

Sandia National Laboratories

ATTN: Tech Lib, 3141

ATTN: 0333, R. Stratton

ATTN: 0334, J. Struve

OTHER GOVERNMENT AGENCIES

Bureau of Alcohol, Tobacco & Firearms

ATTN: Chief Special Opns Div

US Dept of State, Bureau of Politico Mil Affairs

ATTN: PM/STM

Committee on Armed Services

ATTN: Staff Dir & Chief Counsel

OTHER GOVERNMENT AGENCIES (Continued)

Central Intell Agency

ATTN: Counter-Terrorist Group

ATTN: Dir of Security

ATTN: Medical Svcs

ATTN: NIO-T

ATTN: NIO, Strategic Sys

ATTN: Ofc of Global Issues

ATTN: R&D Sub Committee

ATTN: Security Committee

ATTN: Tech Library

Federal Aviation Admin

ATTN: Dir of Civil Aviation Security

Federal Bureau of Invest Academy

ATTN: Behavioral Rsch Unit

2 cys ATTN: Library

Federal Bureau of Investigation

3 cys ATTN: Terrorist Rsch & Analytical Ctr

Federal Emergency Management Agency

ATTN: Asst Assoc Dir for Rsch, J. Kerr

ATTN: Civil Security Division

ATTN: G. Orrell, MP-CP

ATTN: Ofc of Rsch/NP, D. Benson

General Svcs Administration

ATTN: PS

House Perm Select Committ on Intell

ATTN: Staff Director

Interpol, US Natl Central Bureau

ATTN: Chief

Metro Transit Police

ATTN: Chief

National Bureau of Standards

ATTN: Law Enforcement

Dept of Commerce, Natl Bureau of Standards

ATTN: Tech A219

Natl Criminal Justice Reference Svc

2 cys ATTN: D. Galarraga

Select Committee on Intelligence

ATTN: Staff Director

Subcommittee on Sec & Terrorism

ATTN: Chief Counsel, Staff Dir

US Capitol Police

ATTN: Chief

US Coast Guard

ATTN: Port & Environment Safety

US Coast Guard Academy

ATTN: Library

US Dept of State

ATTN: A/SY/CC/TAG

ATTN: A/SY/DASS

ATTN: A/SY/OP/T

ATTN: FAIM/LR

ATTN: M/MED

2 cys ATTN: M/CTP

OTHER GOVERNMENT AGENCIES (Continued)

US Nuclear Regulatory Commission
ATTN: Dir Div of Safeguards
ATTN: Ofc of Insp & Enforcement

US Park Police
ATTN: Chief of Police

DEPARTMENT OF DEFENSE CONTRACTORS

Advanced International Studies Institute
ATTN: M. Harvey

Advanced Rsch & Applications Corp
ATTN: Doc Control

Aerospace Corp
ATTN: Library

Analytical Assessments Corp
ATTN: A. Wagner

BDM Corp

ATTN: C. Wasaff

ATTN: J. Bode

ATTN: J. Braddock

ATTN: J. Conant

ATTN: R. Buchanan

Boeing Co
ATTN: MS-85-20, D. Choate
ATTN: MS-85-20, J. Russel

Computer Sciences Corp
ATTN: F. Eisenbarth

Data Memory Systems, Inc
ATTN: T. Dupuy

Grumman-CITC, Inc
ATTN: S. Shrier

Horizons Technology, Inc
ATTN: J. Palmer

IIT Research Institute
ATTN: Doc Library

Institute for Defense Analyses
ATTN: Classified Library
ATTN: J. Grote

IRT Corp
ATTN: W. Macklin

JAYCOR
ATTN: R. Sullivan

Kaman Sciences Corp
ATTN: F. Shelton

Kaman Sciences Corp
ATTN: E. Conrad
ATTN: E. Daugs

Kaman Tempo
ATTN: DASIAC

Kaman Tempo
ATTN: DASIAC

Kaman Tempo
ATTN: DASIAC

DEPARTMENT OF DEFENSE CONTRACTORS (Continued)

Martin Marietta Corp
ATTN: F. Marion

Martin Marietta Denver Aerospace
ATTN: J. Donathan

National Institute for Public Policy
ATTN: C. Gray

Orion Research Inc
ATTN: J. Scholz

Pacific-Sierra Research Corp
ATTN: H. Brode, Chairman SAGE
ATTN: G. Anno
ATTN: S. Baum
ATTN: R. Young
ATTN: H. Withers

2 cys ATTN: R. Young
2 cys ATTN: S. Baum
2 cys ATTN: H. Withers

Pacific-Sierra Research Corp
ATTN: D. Gormley

R&D Associates

ATTN: C. Lee

ATTN: C. Knowles

ATTN: D. Simons

ATTN: E. Carson

ATTN: F. Field

ATTN: P. Haas

2 cys ATTN: Doc Control

R&D Associates

ATTN: A. Deverill

ATTN: J. Thompson

ATTN: K. Moran

ATTN: W. Graham

Rand Corp

ATTN: P. Davis

ATTN: V. Jackson

2 cys ATTN: Security & Subnation Conflict

Rand Corp

ATTN: B. Bennett

Rockwell International Corp
ATTN: J. Howe

S-CUBED

ATTN: K. Pyatt

Science Applications Intl Corp
ATTN: Document Control
ATTN: E. Swick
ATTN: J. Beyster
ATTN: J. Martin
ATTN: J. Warner
ATTN: M. Drake

Science Applications Intl Corp
ATTN: B. Bennett
ATTN: Document Control
ATTN: J. Foster
ATTN: J. Peters
ATTN: J. Shannon
ATTN: L. Goure
ATTN: M. Fineberg
ATTN: W. Layson

Science Applications Intl Corp
ATTN: D. Kaul

DEPARTMENT OF DEFENSE CONTRACTORS (Continued)

Science Applications, Inc
ATTN: R. Craver

SRI International
ATTN: R. Tidwell

SRI International
ATTN: C. Hulburt

Systems Research & Applications Corp
ATTN: S. Greenstein

DEPARTMENT OF DEFENSE CONTRACTORS (Continued)

Tetra Tech, Inc
ATTN: F. Bothwell

TRW Electronics & Defense Sector
ATTN: D. Scally
ATTN: N. Lipner
ATTN: R. Burnett

TRW Electronics & Defense Sector
ATTN: P. Dai